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(71) Applicant (for all designated States except US): MIL-LENNIUM PHARMACEUTICALS, INC. [US/US]; 75 Sidney Street, Cambridge, MA 02139 (US).

(72) Inventors; and
(75) Inventors/Applicants (for US only): SCHLEGEL, Robert [US/US]; 211 Melrose Street, Auburndale, MA 02466 (US). CHEN, Yan [CN/US]; 26A Plymouth Street, Apartment 2, Cambridge, MA 02141 (US). ZHAO, Xumei [US/US]; 6 Wildwood Lane, Burlington, MA 01803 (US). MONAHAN, John, E. [US/US]; 942 West Street, Walpole, MA 02081 (US). KAMATKAR, Shub-hangi [IN/US]; 655 Saw Mill Brook Parkway, #1, Newton, MA 02459 (US). GANNAVARAPU, Manjula [IN/US]; 10 Windemere Drive, Acton, MA 01720 (US). GLATT,

Karen [US/US]; 17 Beacon Street, Natick, MA 01760 (US). HOERSCH, Sebastian [DE/US]; 127 Brattle Street, Arlington, MA 02424 (US).

(74) Agents: SMITH, DeAnn, F. et al.; Lahive & Cockfield, LLP, 28 State Street, Boston, MA 02109 (US).

(54) Title: NOVEL GENES, COMPOSITIONS, KITS, AND METHODS FOR IDENTIFICATION, ASSESSMENT, PREVEN-TION, AND THERAPY OF CERVICAL CANCER

(57) Abstract: The invention relates to newly discovered nucleic acid molecules and proteins associated with cervical cancer in-cluding pre-malignant conditions such as dysplasia. Compositions, kits, and methods for detecting, characterizing, preventing, and treating human cervical cancers are provided.

NOVEL GENES, COMPOSITIONS, KITS, AND METHODS FOR IDENTIFICATION, ASSESSMENT, PREVENTION, AND THERAPY OF CERVICAL CANCER

5   RELATED APPLICATIONS

The present application claims priority to U.S. provisional patent application serial no. 60/298,159, filed on June 13, 2001, U.S. provisional patent application serial no. 60/298,155, filed on June 13, 2001, and U.S. provisional patent application serial no. 60/335,936, filed on November 14, 2001, all of which are expressly incorporated by

10   reference.

FIELD OF THE INVENTION

The field of the invention is cervical cancer, including diagnosis, characterization, management, and therapy of cervical cancer.

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BACKGROUND OF THE INVENTION

The increased number of cancer cases reported in the United States, and, indeed, around the world, is a major concern. Currently there are only a handful of treatments available for specific types of cancer, and these provide no absolute guarantee

20   of success. In order to be most effective, these treatments require not only an early detection of the malignancy, but a reliable assessment of the severity of the malignancy.

Cancer of the cervix is one of the most common malignancies in women and remains a significant public health problem throughout the world. In the United States alone, invasive cervical cancer accounts for approximately 19% of all

25   gynecological cancers. In 1996, it was estimated that there were 14,700 newly diagnosed cases and 4900 deaths attributed to this disease (American Cancer Society, Cancer Facts & Figures 1996, Atlanta, Ga.: American Cancer Society, 1996). In many developing countries, where mass screening programs are not widely available, the clinical problem is more serious. Worldwide, the number of new cases is estimated to be 471,000 with a

30   four-year survival rate of only 40% (Munoz et al., 1989, *Epidemiology of Cervical Cancer* In: "Human Papillomavirus", New York, Oxford Press, pp 9-39; National Institutes of Health, Consensus Development Conference Statement on Cervical Cancer, Apr.1-3, 1996).

The precursor to cervical cancer is dysplasia, also known in the art as cervical intraepithelial neoplasia (CIN) or squamous intraepithelial lesions (SIL). While it is not understood how normal cells become transformed, the concept of a continuous spectrum of histopathological change from normal, stratified epithelium through CIN to

5   invasive cancer has been widely accepted for many years. A large body of epidemiological and molecular biological evidence has established human papillomavirus (HPV) infection as a causative factor in cervical cancer. HPV is found in 85% or more of squamous cell invasive lesions, which represent the most common histologic type seen in cervical carcinoma. Additional cofactors have also been

10   identified, including oncogenes that have been activated by point mutations and chromosomal translocations or deletions.

In light of this, cervical cancer remains a highly preventable form of cancer when pre-invasive lesions are detected early. Cytological examination of Papanicolaou-stained cervical smears (also referred to as Pap smears) is currently the

15   principle method for detecting cervical cancer. Not surprisingly, the effectiveness of Pap smear screening varies depending not only upon the quality of the sample being used, but also upon subjective parameters that are inherent to the analysis. In addition, despite the historical success of the test, concerns have arisen regarding its ability to reliably predict the behavior of some pre-invasive lesions (Ostor *et al.*, 1993, *Int. J. Gynecol.*

20   *Pathol.* 12: 186-192; and Genest *et al.*, 1993, *Human Pathol.* 24: 730-736).

SUMMARY OF THE INVENTION

The invention relates to cancer markers (hereinafter "markers" or "markers of the inventions"), which are listed in Table 1. The invention provides

25   nucleic acids and proteins that are encoded by or correspond to the markers (hereinafter "marker nucleic acids" and "marker proteins," respectively). Table 1 provides the sequence identifiers of the sequences of such marker nucleic acids and proteins listed in the accompanying Sequence Listing. The invention further provides antibodies, antibody derivatives and antibody fragments which bind specifically with such proteins

30   and/or fragments of the proteins.

The invention also relates to various methods, reagents and kits for diagnosing, staging, prognosing, monitoring and treating cervical cancer. "Cervical cancer " as used herein includes carcinomas, (*e.g.*, carcinoma in situ, invasive

carcinoma, metastatic carcinoma) and pre-malignant conditions, (*e.g.*, dysplasia, including CIN or SIL). In one embodiment, the invention provides a diagnostic method of assessing whether a patient has cervical cancer or has higher than normal risk for developing cervical cancer, comprising the steps of comparing the level of expression of

5 a marker of the invention in a patient sample and the normal level of expression of the marker in a control, *e.g.*, a sample from a patient without cervical cancer. A significantly higher level of expression of the marker in the patient sample as compared to the normal level is an indication that the patient is afflicted with cervical cancer or has higher than normal risk for developing cervical cancer.

10   According to the invention, the markers are selected such that the positive predictive value of the methods of the invention is at least about 10%, preferably about 25%, more preferably about 50% and most preferably about 90%. Also preferred for use in the methods of the invention are markers that are differentially expressed, as compared to normal cervical cells, by at least two-fold in at least about 20%,more

15 preferably about 50% and most preferably about 75% of any of the following conditions: stage 0 cervical cancer patients, stage I cervical cancer patients, stage II cervical cancer patients, stage III cervical cancer patients, stage IV cervical cancer patients, grade I cervical cancer patients, grade II cervical cancer patients, grade III cervical cancer patients, squamous cell (epidermoid) cervical cancer patients, cervical adenocarcinoma

20 patients, cervical adenosquamous carcinoma patients, small-cell cervical carcinoma patients, malignant cervical cancer patients, patients with primary carcinomas of the cervix, patients with primary malignant lymphomas of the cervix and patients with secondary malignant lymphomas of the cervix, and all other types of cancers, malignancies and transformations associated with the cervix.

25   In a preferred diagnostic method of assessing whether a patient is afflicted with cervical cancer (*e.g.*, new detection ("screening"), detection of recurrence, reflex testing), the method comprises comparing:

        a) the level of expression of a marker of the invention in a patient sample, and

30   b) the normal level of expression of the marker in a control non-cervical cancer sample.

A significantly higher level of expression of the marker in the patient sample as compared to the normal level is an indication that the patient is afflicted with cervical cancer.

The invention also provides diagnostic methods for assessing the efficacy of a therapy for inhibiting cervical cancer in a patient. Such methods comprise comparing:

    a) expression of a marker of the invention in a first sample obtained from the patient prior to providing at least a portion of the therapy to the patient, and

    b) expression of the marker in a second sample obtained from the patient following provision of the portion of the therapy.

A significantly lower level of expression of the marker in the second sample relative to that in the first sample is an indication that the therapy is efficacious for inhibiting cervical cancer in the patient.

It will be appreciated that in these methods the "therapy" may be any therapy for treating cervical cancer including, but not limited to, chemotherapy, radiation therapy, surgical removal of tumor tissue, gene therapy and biologic therapy such as the administering of antibodies and chemokines. Thus, the methods of the invention may be used to evaluate a patient before, during and after therapy, for example, to evaluate the reduction in tumor burden.

In a preferred embodiment, the diagnostic methods are directed to therapy using a chemical or biologic agent. These methods comprise comparing:

    a) expression of a marker of the invention in a first sample obtained from the patient and maintained in the presence of the chemical or biologic agent, and

    b) expression of the marker in a second sample obtained from the patient and maintained in the absence of the agent.

A significantly lower level of expression of the marker in the second sample relative to that in the first sample is an indication that the agent is efficacious for inhibiting cervical cancer, in the patient. In one embodiment, the first and second samples can be portions of a single sample obtained from the patient or portions of pooled samples obtained from the patient.

The invention additionally provides a monitoring method for assessing the progression of cervical cancer in a patient, the method comprising:

a) detecting in a patient sample at a first time point, the expression of a marker of the invention;

b) repeating step a) at a subsequent time point in time; and

c) comparing the level of expression detected in steps a) and b), and therefrom monitoring the progression of cervical cancer in the patient.

A significantly higher level of expression of the marker in the sample at the subsequent time point from that of the sample at the first time point is an indication that the cervical cancer has progressed, whereas a significantly lower level of expression is an indication that the cervical cancer has regressed.

The invention further provides a diagnostic method for determining whether cervical cancer has metastasized or is likely to metastasize in the future, the method comprising comparing:

a) the level of expression of a marker of the invention in a patient sample, and

b) the normal level (or non-metastatic level) of expression of the marker in a control sample.

A significantly higher level of expression in the patient sample as compared to the normal level (or non-metastatic level) is an indication that the cervical cancer has metastasized or is likely to metastasize in the future.

The invention moreover provides a test method for selecting a composition for inhibiting cervical cancer in a patient. This method comprises the steps of:

a) obtaining a sample comprising cancer cells from the patient;

b) separately maintaining aliquots of the sample in the presence of a plurality of test compositions;

c) comparing expression of a marker of the invention in each of the aliquots; and

d) selecting one of the test compositions which significantly reduces the level of expression of the marker in the aliquot containing that test composition, relative to the levels of expression of the marker in the presence of the other test compositions.

The invention additionally provides a test method of assessing the cervical carcinogenic potential of a compound. This method comprises the steps of:

a) maintaining separate aliquots of cervical cells in the presence and absence of the compound; and

5 b) comparing expression of a marker of the invention in each of the aliquots.

A significantly higher level of expression of the marker in the aliquot maintained in the presence of the compound, relative to that of the aliquot maintained in the absence of the compound, is an indication that the compound possesses cervical carcinogenic potential.

10 In addition, the invention further provides a method of inhibiting cervical cancer in a patient. This method comprises the steps of:

a) obtaining a sample comprising cancer cells from the patient;

b) separately maintaining aliquots of the sample in the presence of a plurality of compositions;

15 c) comparing expression of a marker of the invention in each of the aliquots; and

d) administering to the patient at least one of the compositions which significantly lowers the level of expression of the marker in the aliquot containing that composition, relative to the levels of expression of the

20 marker in the presence of the other compositions.

In the aforementioned methods, the samples or patient samples comprise cells obtained from the patient. The cells may be found in a cervical smear collected, for example, by a cervical brush. In another embodiment, the sample is a body fluid. Such fluids include, for example, blood fluids, lymph, ascitic fluids, gynecological fluids,

25 urine, and fluids collected by vaginal rinsing. In a further embodiment, the patient sample is *in vivo*.

According to the invention, the level of expression of a marker of the invention in a sample can be assessed, for example, by detecting the presence in the sample of:

30 • the corresponding marker protein (*e.g.*, a protein having one of the sequences set forth as "SEQ ID NO (AAs)" in Table 1, or a fragment of the protein (*e.g.* by using a reagent, such as an antibody, an antibody derivative,

an antibody fragment or single-chain antibody, which binds specifically with
the protein or protein fragment)

- the corresponding marker nucleic acid (*e.g.* a nucleotide transcript having
  one of the nucleic acid sequences set forth as "SEQ ID NO (nts)" in Table 1,
  or a complement thereof), or a fragment of the nucleic acid (*e.g.* by
  contacting transcribed polynucleotides obtained from the sample with a
  substrate having affixed thereto one or more nucleic acids having the entire
  or a segment of the nucleic acid sequence of any of the SEQ ID NO (nts), or
  a complement thereof)

- a metabolite which is produced directly (*i.e.*, catalyzed) or indirectly by
  the corresponding marker protein.

According to the invention, any of the aforementioned methods may be
performed using a plurality (*e.g.* 2, 3, 5, or 10 or more) of cervical cancer markers,
including cervical cancer markers known in the art. In such methods, the level of
expression in the sample of each of a plurality of markers, at least one of which is a
marker of the invention, is compared with the normal level of expression of each of the
plurality of markers in samples of the same type obtained from control humans not
afflicted with cervical cancer. A significantly altered (*i.e.*, increased or decreased as
specified in the above-described methods using a single marker) level of expression in
the sample of one or more markers of the invention, or some combination thereof,
relative to that marker's corresponding normal or control level, is an indication that the
patient is afflicted with cervical cancer. For all of the aforementioned methods, the
marker(s) are preferably selected such that the positive predictive value of the method is
at least about 10%.

In a further aspect, the invention provides an antibody, an antibody
derivative, or an antibody fragment, which binds specifically with a marker protein (*e.g.*,
a protein having one of the amino acid sequences set forth in the Sequence Listing) or a
fragment of the protein. The invention also provides methods for making such antibody,
antibody derivative, and antibody fragment. Such methods may comprise immunizing a
mammal with a protein or peptide comprising the entirety, or a segment of 10 or more
amino acids, of a marker protein (*e.g.*, a protein having one of the amino acid sequences
set forth in the Sequence Listing), wherein the protein or peptide may be obtained from a
cell or by chemical synthesis. The methods of the invention also encompass producing

monoclonal and single-chain antibodies, which would further comprise isolating splenocytes from the immunized mammal, fusing the isolated splenocytes with an immortalized cell line to form hybridomas, and screening individual hybridomas for those that produce an antibody that binds specifically with a marker protein or a

5   fragment of the protein.

In another aspect, the invention relates to various diagnostic and test kits. In one embodiment, the invention provides a kit for assessing whether a patient is afflicted with cervical cancer. The kit comprises a reagent for assessing expression of a marker of the invention. In another embodiment, the invention provides a kit for

10   assessing the suitability of a chemical or biologic agent for inhibiting cervical cancer in a patient. Such a kit comprises a reagent for assessing expression of a marker of the invention, and may also comprise one or more of such agents. In a further embodiment, the invention provides kits for assessing the presence of cervical cancer cells or treating cervical cancers. Such kits comprise an antibody, an antibody derivative, or an antibody

15   fragment, which binds specifically with a marker protein, or a fragment of the protein. Such kits may also comprise a plurality of antibodies, antibody derivatives, or antibody fragments wherein the plurality of such antibody agents binds specifically with a marker protein, or a fragment of the protein.

In an additional embodiment, the invention also provides a kit for

20   assessing the presence of cervical cancer cells, wherein the kit comprises a nucleic acid probe that binds specifically with a marker nucleic acid or a fragment of the nucleic acid. The kit may also comprise a plurality of probes, wherein each of the probes binds specifically with a marker nucleic acid, or a fragment of the nucleic acid.

In a further aspect, the invention relates to methods for treating a patient

25   afflicted with cervical cancer or at risk of developing cervical cancer. Such methods may comprise reducing the expression and/or interfering with the biological function of a marker of the invention. In one embodiment, the method comprises providing to the patient an antisense oligonucleotide or polynucleotide complementary to a marker nucleic acid, or a segment thereof. For example, an antisense polynucleotide may be

30   provided to the patient through the delivery of a vector that expresses an anti-sense polynucleotide of a marker nucleic acid or a fragment thereof. In another embodiment, the method comprises providing to the patient an antibody, an antibody derivative, or antibody fragment, which binds specifically with a marker protein or a fragment of the

protein. In a preferred embodiment, the antibody, antibody derivative or antibody fragment binds specifically with a protein having one of the amino acid sequences set forth in the Sequence Listing, or a fragment of the protein.

It will be appreciated that the methods and kits of the present invention 5 may also include known cancer markers including known cervical cancer markers. It will further be appreciated that the methods and kits may be used to identify cancers other than cervical cancer.

DETAILED DESCRIPTION OF THE INVENTION

10 The invention relates to newly discovered cancer markers associated with the cancerous state of cervical cells. It has been discovered that the higher than normal level of expression of any of these markers or combination of these markers correlates with the presence of cervical cancer including pre-malignant conditions such as dysplasia, in a patient. Methods are provided for detecting the presence of cervical 15 cancer in a sample, the absence of cervical cancer in a sample, the stage of a cervical cancer, and other characteristics of cervical cancer that are relevant to prevention, diagnosis, characterization, and therapy of cervical cancer in a patient. Methods of treating cervical cancer are also provided.

Table 1 lists the markers of the invention which are over-expressed in 20 cervical cancer cells compared to normal (*i.e.*, non-cancerous) cervical cells and comprises markers listed in Tables 2 and 3. Table 2 lists newly-identified nucleotide and amino acid sequences. Table 3 lists newly-identified nucleotide sequences. Tables 1-3 provide the sequence listing identifiers of the cDNA sequence of a nucleotide transcript and the amino acid sequence of a protein encoded by or corresponding to each 25 marker, as well as the location of the protein coding sequence within the cDNA sequence.

Table 1

Marker	Gene Name	SEQ ID NO (nts)	SEQ ID NO (AAs)	CDS
M661	AKAP9: A kinase (PRKA) anchor protein (yotiao) 9, variant 1	1	2	223..11946
M662	AKAP9: A kinase (PRKA) anchor protein (yotiao) 9, variant 2	3	4	223..11922
M663	AKAP9: A kinase (PRKA) anchor protein (yotiao) 9, variant 3	5	6	223..12000
M664	AKAP9: A kinase (PRKA) anchor protein (yotiao) 9, variant 4	7	8	223..11976
M1	APOL1: Apolipoprotein L-I mNA, splice variant A, major form	9	10	213..1364
M2	APOL1: Apolipoprotein L-I mNA, splice variant B, minor form	11	12	274..1518
M3	APOL3: apolipoprotein L, 3; TNF-inducible protein CG12-1	13	14	418..1413
OV3	AQP5: Aquaporin 5	15	16	519..1316
M4	BC001980: clone MGC:5618	17	18	157..225
M5	BST2: Bone marrow stromal cell antigen 2	19	20	10..552
M6	BTEB1: basic transcription element binding protein 1	21	22	1265..1999
M665	CD74: CD74 antigen (invariant polypeptide of major histocompatibility complex,class II antigen-associated)	23	24	8..706
M7	CDC20: CDC20 cell cycle protein	25	26	45..1544
M8	CDKN2C: cyclin-dependent kinase inhibitor 2C, p18	27	28	1216..1722
M9	CKTSF1B1: (cysteine knot superfamily 1, BMP antagonist 1), gremlin	29	30	45..1544
M10	CLDN1: claudin 1	31	32	221..856
M11	CLIC4: chloride intracellular channel 4	33	34	198..959
M12	COL1A1: collagen, type I, alpha 1	35	36	120..4514
M13	COL1A2: collagen, type I, alpha 2	37	38	140..4240
M14	COL8A1: collagen, type VIII, alpha 1	39	40	1..2235
M15	COPA: coatomer protein complex, subunit alpha	41	42	467..4141
M16	CRIP1: cysteine-rich protein 1 (intestinal)	43	44	1..234
M17	CTGF: connective tissue growth factor	45	46	146..1195
M18	DOC: downregulated in ovarian cancer 1	47	48	135..2393
M19	EFNA1: ephrin-A1	49	50	74..691
M481	EPPK1: epiplakin 1	51	52	89..15286
M20	FLJ11350: hypothetical protein FLJ11350	53	54	106..1047
M21	FLJ13809: hypothetical protein FLJ13809	55	56	64..1593
M22	FLJ20500: hypothetical protein FLJ20500	57	58	198..896
M23	FLJ23399: hypothetical protein FLJ23399	59	60	283..1770
M24	FN1: Fibronectin 1, variant 1	61	62	<1..2384
M25	FN1: Fibronectin 1, variant 2	63	64	<1..6988
M482	FOSL2: FOS-like antigen 2, variant 1	65	66	324..1304
M483	FOSL2: FOS-like antigen 2, variant 2	67	66	324..1304
M484	FSHPRH1: FSH primary response (LRPR1, rat) homolog 1	68	69	270..2540
M26	FY: Duffy blood group	70	71	495..1511

M485	G1P3:interferon, alpha-inducible protein (clone IFI-6-16)	72	73	108..500
M486	GW112: GW112 protein	74	75	509..1072
M27	HSKERUV: clone 266, Human radiated keratinocyte mRNA 266 (keratin-related protein)	76	77	<1..801
M28	HSPC121: butyrate-induced transcript 1	78	79	150..1271
M29	HUMCLPB: Coactosin like protein	80	81	150..576
M487	hypothetical protein	82	83	58..8163
M30	IFI27: (interferon, alpha-inducible protein 27	84	85	55..423
OV31	IFI30: interferon, gamma-inducible protein 30	86	87	41..952
M31	IFITM2: interferon induced transmembrane protein 2 (1-8D)	88	89	280..678
M32	IGFBP-3: insulin-like growth factor binding protein 3	90	91	133..1009
M33	IL8RA: interleukin 8	92	93	75..374
M34	INHBA: Inhibin, beta-1	94	95	86..1366
M488	ITGA3: integrin, alpha 3 (antigen CD49C, alpha 3 subunit of VLA-3 receptor), variant a	96	97	74..3229
M454	ITGA3: integrin, alpha 3 (antigen CD49C, alpha 3 subunit of VLA-3 receptor), variant b	98	99	74..3274
M35	ITGB6: integrin, beta 6	100	101	195..2561
M36	KATII: L-kynurenine/alpha-aminoadipate aminotransferase	102	103	454..1731
M666	KCNAB1: potassium voltage-gated channel, shaker-related subfamily, beta member 1, variant 1	104	105	89..1315
M667	KCNAB1: potassium voltage-gated channel, shaker-related subfamily, beta member 1, variant 2	106	107	54..1313
M668	KCNAB1: potassium voltage-gated channel, shaker-related subfamily, beta member 1, variant 3	108	109	28..1233
M37	KIAA0662: KIAA0662 protein	110	111	<1..2035
M38	LAMA3: Laminin, alpha-3 (nicein (150kD), (kalinin (165kD), BM600 (150kD)	112	113	1..5142
M39	LAMC2: laminin, gamma 2	114	115	90..3671
M40	LSM5: U6 snRNA-associated Sm-like protein	116	117	1..276
M41	LUM: lumican	118	119	85..1101
M42	MACMARCKS: macrophage myristoylated alanine-rich C kinase substrate	120	121	14..601
M43	MAGP: microfibrillar-associated protein 2 precursor, transcript variant 1	122	123	115..666
M44	MAGP: microfibrillar-associated protein 2 precursor, transcript variant 2	124	125	100..651
M45	MAPK: mitogen-activated protein kinase 1	126	127	328..1410
M489	MCM6: minichromosome maintenance deficient (mis5, S. pombe) 6	128	129	62..2527
M46	MDK: midkine (neurite growth-promoting factor 2)	130	131	26..457
M47	MGP: matrix Gla protein	132	133	47..358
M48	MMP12: matrix metalloproteinase 12	134	135	13..1425
M49	MMP3: matrix metalloproteinase 3, stromelysin 1, progelatinase	136	137	64..1497
M294	MMP7: matrix metalloproteinase 7 (matrilysin, uterine), PUMP1 proteinase, variant 1	138	139	48..851
OV52	MMP7: matrix metalloproteinase 7 (matrilysin, uterine), PUMP1 proteinase, variant 2	140	139	28..831

M50	MMP9: matrix metalloproteinase 9, gelatinase B, 92kD gelatinase, 92kD type IV collagenase	141	142	20..2143
OV68	MSLN: mesothelin, variant 1	143	144	88..2196
OV69	MSLN: mesothelin, variant 2	145	146	88..1980
OV70	MSLN: mesothelin, variant 3	147	148	88..1950
OV71	MSLN: mesothelin, variant 4	149	150	88..2172
OV72	MSLN: mesothelin, variant 5	151	152	88..1926
OV43	MSLN: mesothelin, variant 6	153	154	88..1956
OV45	MUC1: mucin 1, transmembrane, variant 1	155	156	58..1605
M669	MUC1: mucin 1, transmembrane, variant 2	157	158	74..3841
M51	MYBL2: v-myb avian myeloblastosis viral oncogene homolog-like 2	159	160	128..2230
M52	MYH11: smooth muscle myosin heavy chain 11, isoform SM1	161	162	89..6007
M53	MYH11: smooth muscle myosin heavy chain 11, isoform SM2	163	164	89..5905
M54	NK4: natural killer cell transcript 4 , variant 1	165	166	60..764
M670	NK4: natural killer cell transcript 4 , variant 2	167	168	60..764
M55	NP25: (neuronal protein)	169	170	50..898
OV48	OPN-a (osteopontin), SPP1 (secreted phosphoprotein 1), bone sialoprotein I	171	172	1..942
OV49	OPN-b (osteopontin), SPP1 (secreted phosphoprotein 1), bone sialoprotein I	173	174	88..990
OV50	OPN-c (osteopontin), SPP1 (secreted phosphoprotein 1), bone sialoprotein I	175	176	1..861
M56	OSF-2, osteoblast specific factor 2 (fasciclin I-like), variant 1	177	178	12..2522
M491	OSF-2, osteoblast specific factor 2 (fasciclin I-like), variant 2	179	180	28..2367
M57	PIM2: pim-2 oncogene	181	182	186..1190
M58	PLAU: plasminogen activator, urokinase	183	184	77..1372
M59	PLK: polo (Drosophia)-like kinase	185	186	64..1875
M671	PNN: pinin, desmosome associated protein	187	188	31..2262
M60	PRG1: proteoglycan 1, secretory granule	189	190	25..501
M61	PTHLH: parathyroid hormone-like hormone	191	192	304..831
M62	PTN: pleiotrophin (heparin binding growth factor 8, neurite growth-promoting factor 1)	193	194	1542..2048
M63	RAB6KIFL: RAB6 interacting, kinesin-like (rabkinesin6)	195	196	28..2700
M64	RARRES3: retinoic acid receptor responder (tazarotene induced) 3	197	198	62..556
M65	RBP1: retinol-binding protein 1(cellular), CRABP-I, CRBP-I	199	200	126..533
M66	RGS16: Regulator of G protein signaling-16	201	202	93..701
M67	S100A2: S100 calcium binding protein A2, variant 1	203	204	72..362
M68	S100A2: S100 calcium binding protein A2, variant 2	205	206	41..334
M69	SCYA20: small inducible cytokine subfamily A (Cys-Cys), member 20	207	208	59..349
M70	SPARC: Osteonectin (secreted protein, acidic, cysteine-rich)	209	210	58..969
M71	STCH: stress 70 protein chaperone, microsome-associated	211	212	37..1452
M492	STK12: serine/ threonine kinase 12	213	214	58..1092

M72	TK1: thymidine kinase 1, soluble	215	216	58..762
OV86	TMPRSS4: transmembrane protease, serine 4	217	218	310..1623
M73	TMSB4X: thymosin, beta 4, X chromosome	219	220	78..212
M74	TOP2A: topoisomerase (DNA) II alpha (170kD)	221	222	37..4632
M493	TPM1: tropomyosin 1 (alpha)	223	224	57..911
M75	TXN: thioredoxin	225	226	64..381
M76	UBCH10: ubiquitin carrier protein E2-C	227	228	41..580
M77	UBD: diubiquitin	229	230	19..516
M78	unnamed gene (1)	231	232	45..1353
M79	unnamed gene (2)	233	234	1..1508
M80	VATD: vacuolar proton pump delta polypeptide	235	236	166..909
M81	ZWINT: ZW10 interactor	237	238	25..858

Table 2

Marker	Gene Name	SEQ ID NO (nts)	SEQ ID NO (AAs)	CDS
M661	AKAP9: A kinase (PRKA) anchor protein (yotiao) 9, variant 1	1	2	223..11946
M662	AKAP9: A kinase (PRKA) anchor protein (yotiao) 9, variant 2	3	4	223..11922
M663	AKAP9: A kinase (PRKA) anchor protein (yotiao) 9, variant 3	5	6	223..12000
M664	AKAP9: A kinase (PRKA) anchor protein (yotiao) 9, variant 4	7	8	223..11976
OV68	MSLN: mesothelin, variant 1	143	144	88..2196
OV69	MSLN: mesothelin, variant 2	145	146	88..1980
OV70	MSLN: mesothelin, variant 3	147	148	88..1950
OV71	MSLN: mesothelin, variant 4	149	150	88..2172
OV72	MSLN: mesothelin, variant 5	151	152	88..1926
M670	NK4: natural killer cell transcript 4 , variant 2	167	168	60..764
M67	S100A2: S100 calcium binding protein A2, variant 1	203	204	72..362
OV86	TMPRSS4: transmembrane protease, serine 4	217	218	310..1623
M78	unnamed gene (1)	231	232	45..1353
M79	unnamed gene (2)	233	234	1..1508

Table 3

Marker	Gene Name	SEQ ID NO (nts)	SEQ ID NO (AAs)	CDS
M481	EPPK1: epiplakin 1	51	52	89..15286
M482	FOSL2: FOS-like antigen 2, variant 1	65	66	324..1304
M483	FOSL2: FOS-like antigen 2, variant 2	67	66	324..1304
M484	FSHPRH1: FSH primary response (LRPR1, rat) homolog 1	68	69	270..2540
M35	ITGB6: integrin, beta 6	100	101	195..2561
OV43	MSLN: mesothelin, variant 6	153	154	88..1956

Definitions

5   As used herein, each of the following terms has the meaning associated with it in this section.

The articles "a" and "an" are used herein to refer to one or to more than one (*i.e.* to at least one) of the grammatical object of the article. By way of example, "an element" means one element or more than one element.

10   A "marker" is a gene whose altered level of expression in a tissue or cell from its expression level in normal or healthy tissue or cell is associated with a disease state, such as cancer. A "marker nucleic acid" is a nucleic acid (*e.g.*, mRNA, cDNA) encoded by or corresponding to a marker of the invention. Such marker nucleic acids include DNA (*e.g.*, cDNA) comprising the entire or a partial sequence of any of the

15   nucleic acid sequences set forth in the Sequence Listing or the complement of such a sequence. The marker nucleic acids also include RNA comprising the entire or a partial sequence of any of the nucleic acid sequences set forth in the Sequence Listing or the complement of such a sequence, wherein all thymidine residues are replaced with uridine residues. A "marker protein" is a protein encoded by or corresponding to a

20   marker of the invention. A marker protein comprises the entire or a partial sequence of any of the sequences set forth in the Sequence Listing. The terms "protein" and "polypeptide' are used interchangeably.

The term "probe" refers to any molecule which is capable of selectively binding to a specifically intended target molecule, for example, a nucleotide transcript or

25   protein encoded by or corresponding to a marker. Probes can be either synthesized by one skilled in the art, or derived from appropriate biological preparations. For purposes of detection of the target molecule, probes may be specifically designed to be labeled, as

described herein. Examples of molecules that can be utilized as probes include, but are not limited to, RNA, DNA, proteins, antibodies, and organic molecules.

A "cervical-associated" body fluid is a fluid which, when in the body of a patient, contacts or passes through cervical cells or into which cells or proteins shed from cervical cells are capable of passing. The cells may be found in a cervical smear collected, for example, by a cervical brush. Exemplary cervical-associated body fluids include blood fluids, lymph, ascitic fluids, gynecological fluids, cystic fluid, urine, and fluids collected by vaginal rinsing.

The "normal" level of expression of a marker is the level of expression of the marker in cervical cells of a human subject or patient not afflicted with cervical cancer

An "over-expression" or "significantly higher level of expression" of a marker refers to an expression level in a test sample that is greater than the standard error of the assay employed to assess expression, and is preferably at least twice, and more preferably three, four, five or ten times the expression level of the marker in a control sample (e.g., sample from a healthy subjects not having the marker associated disease) and preferably, the average expression level of the marker in several control samples.

A "significantly lower level of expression" of a marker refers to an expression level in a test sample that is at least twice, and more preferably three, four, five or ten times lower than the expression level of the marker in a control sample (e.g., sample from a healthy subject not having the marker associated disease) and preferably, the average expression level of the marker in several control samples.

As used herein, the term "promoter/regulatory sequence" means a nucleic acid sequence which is required for expression of a gene product operably linked to the promoter/regulatory sequence. In some instances, this sequence may be the core promoter sequence and in other instances, this sequence may also include an enhancer sequence and other regulatory elements which are required for expression of the gene product. The promoter/regulatory sequence may, for example, be one which expresses the gene product in a tissue-specific manner.

A "constitutive" promoter is a nucleotide sequence which, when operably linked with a polynucleotide which encodes or specifies a gene product, causes the gene product to be produced in a living human cell under most or all physiological conditions of the cell.

An "inducible" promoter is a nucleotide sequence which, when operably linked with a polynucleotide which encodes or specifies a gene product, causes the gene product to be produced in a living human cell substantially only when an inducer which corresponds to the promoter is present in the cell.

A "tissue-specific" promoter is a nucleotide sequence which, when operably linked with a polynucleotide which encodes or specifies a gene product, causes the gene product to be produced in a living human cell substantially only if the cell is a cell of the tissue type corresponding to the promoter.

A "transcribed polynucleotide" or "nucleotide transcript" is a polynucleotide (*e.g.* an mRNA, hnRNA, a cDNA, or an analog of such RNA or cDNA) which is complementary to or homologous with all or a portion of a mature mRNA made by transcription of a marker of the invention and normal post-transcriptional processing (*e.g.* splicing), if any, of the RNA transcript, and reverse transcription of the RNA transcript.

"Complementary" refers to the broad concept of sequence complementarity between regions of two nucleic acid strands or between two regions of the same nucleic acid strand. It is known that an adenine residue of a first nucleic acid region is capable of forming specific hydrogen bonds ("base pairing") with a residue of a second nucleic acid region which is antiparallel to the first region if the residue is thymine or uracil. Similarly, it is known that a cytosine residue of a first nucleic acid strand is capable of base pairing with a residue of a second nucleic acid strand which is antiparallel to the first strand if the residue is guanine. A first region of a nucleic acid is complementary to a second region of the same or a different nucleic acid if, when the two regions are arranged in an antiparallel fashion, at least one nucleotide residue of the first region is capable of base pairing with a residue of the second region. Preferably, the first region comprises a first portion and the second region comprises a second portion, whereby, when the first and second portions are arranged in an antiparallel fashion, at least about 50%, and preferably at least about 75%, at least about 90%, or at least about 95% of the nucleotide residues of the first portion are capable of base pairing

with nucleotide residues in the second portion. More preferably, all nucleotide residues of the first portion are capable of base pairing with nucleotide residues in the second portion.

"Homologous" as used herein, refers to nucleotide sequence similarity between two regions of the same nucleic acid strand or between regions of two different nucleic acid strands. When a nucleotide residue position in both regions is occupied by the same nucleotide residue, then the regions are homologous at that position. A first region is homologous to a second region if at least one nucleotide residue position of each region is occupied by the same residue. Homology between two regions is expressed in terms of the proportion of nucleotide residue positions of the two regions that are occupied by the same nucleotide residue. By way of example, a region having the nucleotide sequence 5'-ATTGCC-3' and a region having the nucleotide sequence 5'-TATGGC-3' share 50% homology. Preferably, the first region comprises a first portion and the second region comprises a second portion, whereby, at least about 50%, and preferably at least about 75%, at least about 90%, or at least about 95% of the nucleotide residue positions of each of the portions are occupied by the same nucleotide residue. More preferably, all nucleotide residue positions of each of the portions are occupied by the same nucleotide residue.

A molecule is "fixed" or "affixed" to a substrate if it is covalently or non-covalently associated with the substrate such the substrate can be rinsed with a fluid (*e.g.* standard saline citrate, pH 7.4) without a substantial fraction of the molecule dissociating from the substrate.

As used herein, a "naturally-occurring" nucleic acid molecule refers to an RNA or DNA molecule having a nucleotide sequence that occurs in an organism found in nature.

A cancer is "inhibited" if at least one symptom of the cancer is alleviated, terminated, slowed, or prevented. As used herein, cervical cancer is also "inhibited" if recurrence or metastasis of the cancer is reduced, slowed, delayed, or prevented.

A kit is any manufacture (*e.g.* a package or container) comprising at least one reagent, *e.g.* a probe, for specifically detecting the expression of a marker of the invention. The kit may be promoted, distributed, or sold as a unit for performing the methods of the present invention.

"Proteins of the invention" encompass marker proteins and their fragments; variant marker proteins and their fragments; peptides and polypeptides comprising an at least 15 amino acid segment of a marker or variant marker protein; and fusion proteins comprising a marker or variant marker protein, or an at least 15 amino acid segment of a marker or variant marker protein.

Unless otherwise specified herewithin, the terms "antibody" and "antibodies" broadly encompass naturally-occurring forms of antibodies (*e.g.*, IgG, IgA, IgM, IgE) and recombinant antibodies such as single-chain antibodies, chimeric and humanized antibodies and multi-specific antibodies, as well as fragments and derivatives of all of the foregoing, which fragments and derivatives have at least an antigenic binding site. Antibody derivatives may comprise a protein or chemical moiety conjugated to an antibody.

Description

The present invention is based, in part, on newly identified markers which are over-expressed in cervical cancer cells as compared to their expression in normal (*i.e.* non-cancerous) cervical cells. The enhanced expression of one or more of these markers in cervical cells is herein correlated with the cancerous state of the tissue. The invention provides compositions, kits, and methods for assessing the cancerous state of cervical cells (*e.g.* cells obtained from a human, cultured human cells, archived or preserved human cells and *in vivo* cells) as well as treating patients afflicted with cervical cancer.

The compositions, kits, and methods of the invention have the following uses, among others:

1) assessing whether a patient is afflicted with cervical cancer;

2) assessing the stage of cervical cancer in a human patient;

3) assessing the grade of cervical cancer in a patient;

4) assessing the benign or malignant nature of cervical cancer in a patient;

5) assessing the metastatic potential of cervical cancer in a patient;

6) assessing the histological type of neoplasm associated with cervical cancer in a patient;

7) making antibodies, antibody fragments or antibody derivatives that are useful for treating cervical cancer and/or assessing whether a patient is afflicted with cervical cancer;

8) assessing the presence of cervical cancer cells;

9) assessing the efficacy of one or more test compounds for inhibiting cervical cancer in a patient;

10) assessing the efficacy of a therapy for inhibiting cervical cancer in a patient;

11) monitoring the progression of cervical cancer in a patient;

12) selecting a composition or therapy for inhibiting cervical cancer in a patient;

13) treating a patient afflicted with cervical cancer;

14) inhibiting cervical cancer in a patient;

15) assessing the cervical carcinogenic potential of a test compound; and

16) preventing the onset of cervical cancer in a patient at risk for developing cervical cancer.

The invention thus includes a method of assessing whether a patient is afflicted with cervical cancer which includes assessing whether the patient has pre-metastasized cervical cancer. This method comprises comparing the level of expression of a marker of the invention (listed in Table 1) in a patient sample and the normal level of expression of the marker in a control, e.g., a non-cervical cancer sample. A significantly higher level of expression of the marker in the patient sample as compared to the normal level is an indication that the patient is afflicted with cervical cancer.

Gene delivery vehicles, host cells and compositions (all described herein) containing nucleic acids comprising the entirety, or a segment of 15 or more nucleotides, of any of the nucleic acid sequences set forth in the Sequence Listing, or the complement of such sequences, and polypeptides comprising the entirety, or a segment of 10 or more amino acids, of any of the amino acid sequences set forth in the Sequence Listing, are also provided by this invention.

As described herein, cervical cancer in patients is associated with an increased level of expression of one or more markers of the invention. While, as discussed above, some of these changes in expression level result from occurrence of the

cervical cancer, others of these changes induce, maintain, and promote the cancerous state of cervical cancer cells. Thus, cervical cancer characterized by an increase in the level of expression of one or more markers of the invention can be inhibited by reducing and/or interfering with the expression of the markers and/or function of the proteins

5   encoded by those markers.

Expression of a marker of the invention can be inhibited in a number of ways generally known in the art. For example, an antisense oligonucleotide can be provided to the cervical cancer cells in order to inhibit transcription, translation, or both, of the marker(s). Alternately, a polynucleotide encoding an antibody, an antibody

10   derivative, or an antibody fragment which specifically binds a marker protein, and operably linked with an appropriate promoter/regulator region, can be provided to the cell in order to generate intracellular antibodies which will inhibit the function or activity of the protein. The expression and/or function of a marker may also be inhibited by treating the cervical cancer cell with an antibody, antibody derivative or antibody

15   fragment that specifically binds a marker protein. Using the methods described herein, a variety of molecules, particularly including molecules sufficiently small that they are able to cross the cell membrane, can be screened in order to identify molecules which inhibit expression of a marker or inhibit the function of a marker protein. The compound so identified can be provided to the patient in order to inhibit cervical cancer

20   cells of the patient.

Any marker or combination of markers of the invention, as well as any known markers in combination with the markers of the invention, may be used in the compositions, kits, and methods of the present invention. In general, it is preferable to use markers for which the difference between the level of expression of the marker in

25   cervical cancer cells and the level of expression of the same marker in normal cervical cells is as great as possible. Although this difference can be as small as the limit of detection of the method for assessing expression of the marker, it is preferred that the difference be at least greater than the standard error of the assessment method, and preferably a difference of at least 2-, 3-, 4-, 5-, 6-, 7-, 8-, 9-, 10-, 15-, 20-, 25-, 100-,

30   500-, 1000-fold or greater than the level of expression of the same marker in normal cervical tissue.

It is recognized that certain marker proteins are secreted from cervical cells (*i.e.* one or both of normal and cancerous cells) to the extracellular space surrounding the cells. These markers are preferably used in certain embodiments of the compositions, kits, and methods of the invention, owing to the fact that the such marker

5   proteins can be detected in a cervical-associated body fluid sample, which may be more easily collected from a human patient than a tissue biopsy sample. In addition, preferred *in vivo* techniques for detection of a marker protein include introducing into a subject a labeled antibody directed against the protein. For example, the antibody can be labeled with a radioactive marker whose presence and location in a subject can be detected by

10   standard imaging techniques.

It is a simple matter for the skilled artisan to determine whether any particular marker protein is a secreted protein. In order to make this determination, the marker protein is expressed in, for example, a mammalian cell, preferably a human cervical cell line, extracellular fluid is collected, and the presence or absence of the

15   protein in the extracellular fluid is assessed (*e.g.* using a labeled antibody which binds specifically with the protein).

The following is an example of a method which can be used to detect secretion of a protein. About 8×10^5 293T cells are incubated at 37°C in wells containing growth medium (Dulbecco's modified Eagle's medium {DMEM}

20   supplemented with 10% fetal bovine serum) under a 5% (v/v) CO_2, 95% air atmosphere to about 60-70% confluence. The cells are then transfected using a standard transfection mixture comprising 2 micrograms of DNA comprising an expression vector encoding the protein and 10 microliters of LipofectAMINE™ (GIBCO/BRL Catalog no. 18342-012) per well. The transfection mixture is maintained for about 5 hours, and then

25   replaced with fresh growth medium and maintained in an air atmosphere. Each well is gently rinsed twice with DMEM which does not contain methionine or cysteine (DMEM-MC; ICN Catalog no. 16-424- 54). About 1 milliliter of DMEM-MC and about 50 microcuries of Trans-^{35}S™ reagent (ICN Catalog no. 51006) are added to each well. The wells are maintained under the 5% CO_2 atmosphere described above and

30   incubated at 37°C for a selected period. Following incubation, 150 microliters of conditioned medium is removed and centrifuged to remove floating cells and debris.

The presence of the protein in the supernatant is an indication that the protein is
secreted.

It will be appreciated that patient samples containing cervical cells may
be used in the methods of the present invention. In these embodiments, the level of
expression of the marker can be assessed by assessing the amount (*e.g.* absolute amount
or concentration) of the marker in a cervical cell sample, *e.g.,* cervical smear obtained
from a patient. The cell sample can, of course, be subjected to a variety of well-known
post-collection preparative and storage techniques (*e.g.,* nucleic acid and/or protein
extraction, fixation, storage, freezing, ultrafiltration, concentration, evaporation,
centrifugation, etc.) prior to assessing the amount of the marker in the sample.
Likewise, cervical smears may also be subjected to post-collection preparative and
storage techniques, *e.g.,* fixation.

The compositions, kits, and methods of the invention can be used to
detect expression of marker proteins having at least one portion which is displayed on
the surface of cells which express it. It is a simple matter for the skilled artisan to
determine whether a marker protein, or a portion thereof, is exposed on the cell surface.
For example, immunological methods may be used to detect such proteins on whole
cells, or well known computer-based sequence analysis methods may be used to predict
the presence of at least one extracellular domain (*i.e.* including both secreted proteins
and proteins having at least one cell-surface domain). Expression of a marker protein
having at least one portion which is displayed on the surface of a cell which expresses it
may be detected without necessarily lysing the cell (*e.g.* using a labeled antibody which
binds specifically with a cell-surface domain of the protein).

Expression of a marker of the invention may be assessed by any of a wide
variety of well known methods for detecting expression of a transcribed nucleic acid or
protein. Non-limiting examples of such methods include immunological methods for
detection of secreted, cell-surface, cytoplasmic, or nuclear proteins, protein purification
methods, protein function or activity assays, nucleic acid hybridization methods, nucleic
acid reverse transcription methods, and nucleic acid amplification methods.

In a preferred embodiment, expression of a marker is assessed using an
antibody (*e.g.* a radio-labeled, chromophore-labeled, fluorophore-labeled, or enzyme-
labeled antibody), an antibody derivative (*e.g.* an antibody conjugated with a substrate or
with the protein or ligand of a protein-ligand pair {*e.g.* biotin-streptavidin}), or an

antibody fragment (*e.g.* a single-chain antibody, an isolated antibody hypervariable domain, etc.) which binds specifically with a marker protein or fragment thereof, including a marker protein which has undergone all or a portion of its normal post-translational modification.

5   In another preferred embodiment, expression of a marker is assessed by preparing mRNA/cDNA (*i.e.* a transcribed polynucleotide) from cells in a patient sample, and by hybridizing the mRNA/cDNA with a reference polynucleotide which is a complement of a marker nucleic acid, or a fragment thereof. cDNA can, optionally, be amplified using any of a variety of polymerase chain reaction methods prior to

10   hybridization with the reference polynucleotide; preferably, it is not amplified. Expression of one or more markers can likewise be detected using quantitative PCR to assess the level of expression of the marker(s). Alternatively, any of the many known methods of detecting mutations or variants (*e.g.* single nucleotide polymorphisms, deletions, etc.) of a marker of the invention may be used to detect occurrence of a

15   marker in a patient.

           In a related embodiment, a mixture of transcribed polynucleotides obtained from the sample is contacted with a substrate having fixed thereto a polynucleotide complementary to or homologous with at least a portion (*e.g.* at least 7, 10, 15, 20, 25, 30, 40, 50, 100, 500, or more nucleotide residues) of a marker nucleic

20   acid. If polynucleotides complementary to or homologous with are differentially detectable on the substrate (*e.g.* detectable using different chromophores or fluorophores, or fixed to different selected positions), then the levels of expression of a plurality of markers can be assessed simultaneously using a single substrate (*e.g.* a "gene chip" microarray of polynucleotides fixed at selected positions). When a method of

25   assessing marker expression is used which involves hybridization of one nucleic acid with another, it is preferred that the hybridization be performed under stringent hybridization conditions.

           Because the compositions, kits, and methods of the invention rely on detection of a difference in expression levels of one or more markers of the invention, it

30   is preferable that the level of expression of the marker is significantly greater than the minimum detection limit of the method used to assess expression in at least one of normal cervical cells and cancerous cervical cells.

It is understood that by routine screening of additional patient samples using one or more of the markers of the invention, it will be realized that certain of the markers are over-expressed in cancers of various types, including specific cervical cancers, as well as other cancers such as breast cancer, ovarian cancer, etc. For
5   example, it will be confirmed that some of the markers of the invention are over-expressed in most (i.e. 50% or more) or substantially all (i.e. 80% or more) of cervical cancer. Furthermore, it will be confirmed that certain of the markers of the invention are associated with cervical cancer of various stages (i.e. stage 0, I, II, III, and IV cervical cancers, as well as subclassifications IA1, IA2, IB, IB1, IB2, IIA, IIB, IIIA, IIIB, IVA,
10   and IVB, using the FIGO Stage Grouping system for primary carcinoma of the cervix (see Gynecologic Oncology, 1991, 41:199 and Cancer, 1992, 69:482)), and pre-malignant conditions (e.g., dysplasia including CIN or SIL), of various histologic subtypes (e.g. squamous cell carcinomas and squamous cell carcinoma variants such as verrucous carcinoma, lymphoepithelioma-like carcinoma, papillary squamous neoplasm
15   and spindle cell squamous cell carcinoma (see Cervical Cancer and Preinvasive Neoplasia, 1996, pp. 90-91) serous, mucinous, endometrioid, and clear cell subtypes, as well as subclassifications and alternate classifications adenocarcinoma, papillary adenocarcinoma, papillary cystadenocarcinoma, surface papillary carcinoma, malignant adenofibroma, cystadenofibroma, adenocarcinoma, cystadenocarcinoma,
20   adenoacanthoma, endometrioid stromal sarcoma, mesodermal {Müllerian} mixed tumor, malignant carcinoma, Brenner tumor, mixed epithelial tumor, and undifferentiated . carcinoma, using the WHO/FIGO system for classification of malignant cervical tumors; Scully, Atlas of Tumor Pathology, 3d series, Washington DC), and various grades (i.e. grade I {well differentiated} , grade II {moderately well differentiated}, and grade III
25   {poorly differentiated from surrounding normal tissue}). In addition, as a greater number of patient samples are assessed for expression of the markers of the invention and the outcomes of the individual patients from whom the samples were obtained are correlated, it will also be confirmed that altered expression of certain of the markers of the invention are strongly correlated with malignant cancers and that altered expression
30   of other markers of the invention are strongly correlated with benign tumors. The compositions, kits, and methods of the invention are thus useful for characterizing one or more of the stage, grade, histological type, and benign/malignant nature of cervical cancer in patients.

When the compositions, kits, and methods of the invention are used for characterizing one or more of the stage, grade, histological type, and benign/malignant nature of cervical cancer in a patient, it is preferred that the marker or panel of markers of the invention is selected such that a positive result is obtained in at least about 20%,

5   and preferably at least about 40%, 60%, or 80%, and more preferably in substantially all patients afflicted with a cervical cancer of the corresponding stage, grade, histological type, or benign/malignant nature. Preferably, the marker or panel of markers of the invention is selected such that a positive predictive value (PPV) of greater than about 10% is obtained for the general population (more preferably coupled with an assay

10   specificity greater than 80%).

When a plurality of markers of the invention are used in the compositions, kits, and methods of the invention, the level of expression of each marker in a patient sample can be compared with the normal level of expression of each of the plurality of markers in non-cancerous samples of the same type, either in a single

15   reaction mixture (*i.e.* using reagents, such as different fluorescent probes, for each marker) or in individual reaction mixtures corresponding to one or more of the markers. In one embodiment, a significantly increased level of expression of more than one of the plurality of markers in the sample, relative to the corresponding normal levels, is an indication that the patient is afflicted with cervical cancer. When a plurality of markers

20   is used, it is preferred that 2, 3, 4, 5, 8, 10, 12, 15, 20, 30, or 50 or more individual markers be used, wherein fewer markers are preferred.

In order to maximize the sensitivity of the compositions, kits, and methods of the invention (*i.e.* by interference attributable to cells of non-cervical origin in a patient sample), it is preferable that the marker of the invention used therein be a

25   marker which has a restricted tissue distribution, *e.g.*, normally not expressed in a non-cervical tissue.

Only a small number of markers are known to be associated with cervical cancer (*e.g.* bcl-2, 15A8 antigen, cdc6, Mcm5, and EGFR). These markers are not, of course, included among the markers of the invention, although they may be used

30   together with one or more markers of the invention in a panel of markers, for example. It is well known that certain types of genes, such as oncogenes, tumor suppressor genes, growth factor-like genes, protease-like genes, and protein kinase-like genes are often involved with development of cancers of various types. Thus, among the markers of the

invention, use of those which correspond to proteins which resemble known proteins encoded by known oncogenes and tumor suppressor genes, and those which correspond to proteins which resemble growth factors, proteases, and protein kinases are preferred.

It is recognized that the compositions, kits, and methods of the invention will be of particular utility to patients having an enhanced risk of developing cervical cancer and their medical advisors. Patients recognized as having an enhanced risk of developing cervical cancer include, for example, patients having a familial history of cervical cancer, patients identified as having a mutant oncogene (*i.e.* at least one allele), and patients of advancing age (*i.e.* women older than about 50 or 60 years).

The level of expression of a marker in normal (*i.e.* non-cancerous) human cervical tissue can be assessed in a variety of ways. In one embodiment, this normal level of expression is assessed by assessing the level of expression of the marker in a portion of cervical cells which appears to be non-cancerous and by comparing this normal level of expression with the level of expression in a portion of the cervical cells which is suspected of being cancerous. Alternately, and particularly as further information becomes available as a result of routine performance of the methods described herein, population-average values for normal expression of the markers of the invention may be used. In other embodiments, the 'normal' level of expression of a marker may be determined by assessing expression of the marker in a patient sample obtained from a non-cancer-afflicted patient, from a patient sample obtained from a patient before the suspected onset of cervical cancer in the patient, from archived patient samples, and the like.

The invention includes compositions, kits, and methods for assessing the presence of cervical cancer cells in a sample (*e.g.* an archived tissue sample or a sample obtained from a patient). These compositions, kits, and methods are substantially the same as those described above, except that, where necessary, the compositions, kits, and methods are adapted for use with samples other than patient samples. For example, when the sample to be used is a parafinized, archived human tissue sample, it can be necessary to adjust the ratio of compounds in the compositions of the invention, in the kits of the invention, or the methods used to assess levels of marker expression in the sample. Such methods are well known in the art and within the skill of the ordinary artisan.

The invention includes a kit for assessing the presence of cervical cancer cells (*e.g.* in a sample such as a patient sample). The kit comprises a plurality of reagents, each of which is capable of binding specifically with a marker nucleic acid or protein. Suitable reagents for binding with a marker protein include antibodies, antibody derivatives, antibody fragments, and the like. Suitable reagents for binding with a marker nucleic acid (*e.g.* a genomic DNA, an mRNA, a spliced mRNA, a cDNA, or the like) include complementary nucleic acids. For example, the nucleic acid reagents may include oligonucleotides (labeled or non-labeled) fixed to a substrate, labeled oligonucleotides not bound with a substrate, pairs of PCR primers, molecular beacon probes, and the like.

The kit of the invention may optionally comprise additional components useful for performing the methods of the invention. By way of example, the kit may comprise fluids (*e.g.* SSC buffer) suitable for annealing complementary nucleic acids or for binding an antibody with a protein with which it specifically binds, one or more sample compartments, an instructional material which describes performance of a method of the invention, a sample of normal cervical cells, a sample of cervical cancer cells, and the like.

The invention also includes a method of making an isolated hybridoma which produces an antibody useful for assessing whether patient is afflicted with an cervical cancer. In this method, a protein or peptide comprising the entirety or a segment of a marker protein is synthesized or isolated (*e.g.* by purification from a cell in which it is expressed or by transcription and translation of a nucleic acid encoding the protein or peptide *in vivo* or *in vitro* using known methods). A vertebrate, preferably a mammal such as a mouse, rat, rabbit, or sheep, is immunized using the protein or peptide. The vertebrate may optionally (and preferably) be immunized at least one additional time with the protein or peptide, so that the vertebrate exhibits a robust immune response to the protein or peptide. Splenocytes are isolated from the immunized vertebrate and fused with an immortalized cell line to form hybridomas, using any of a variety of methods well known in the art. Hybridomas formed in this manner are then screened using standard methods to identify one or more hybridomas which produce an antibody which specifically binds with the marker protein or a fragment thereof. The invention also includes hybridomas made by this method and antibodies made using such hybridomas.

The invention also includes a method of assessing the efficacy of a test compound for inhibiting cervical cancer cells. As described above, differences in the level of expression of the markers of the invention correlate with the cancerous state of cervical cells. Although it is recognized that changes in the levels of expression of

5   certain of the markers of the invention likely result from the cancerous state of cervical cells, it is likewise recognized that changes in the levels of expression of other of the markers of the invention induce, maintain, and promote the cancerous state of those cells. Thus, compounds which inhibit an cervical cancer in a patient will cause the level of expression of one or more of the markers of the invention to change to a level nearer

10   the normal level of expression for that marker (i.e. the level of expression for the marker in non-cancerous cervical cells).

This method thus comprises comparing expression of a marker in a first cervical cell sample and maintained in the presence of the test compound and expression of the marker in a second cervical cell sample and maintained in the absence of the test

15   compound. A significantly reduced expression of a marker of the invention in the presence of the test compound is an indication that the test compound inhibits cervical cancer. The cervical cell samples may, for example, be aliquots of a single sample of normal cervical cells obtained from a patient, pooled samples of normal cervical cells obtained from a patient, cells of a normal cervical cell line, aliquots of a single sample of

20   cervical cancer cells obtained from a patient, pooled samples of cervical cancer cells obtained from a patient, cells of an cervical cancer cell line, or the like. In one embodiment, the samples are cervical cancer cells obtained from a patient and a plurality of compounds known to be effective for inhibiting various cervical cancers are tested in order to identify the compound which is likely to best inhibit the cervical cancer in the

25   patient.

This method may likewise be used to assess the efficacy of a therapy for inhibiting cervical cancer in a patient. In this method, the level of expression of one or more markers of the invention in a pair of samples (one subjected to the therapy, the other not subjected to the therapy) is assessed. As with the method of assessing the

30   efficacy of test compounds, if the therapy induces a significantly lower level of expression of a marker of the invention then the therapy is efficacious for inhibiting cervical cancer. As above, if samples from a selected patient are used in this method,

then alternative therapies can be assessed *in vitro* in order to select a therapy most likely to be efficacious for inhibiting cervical cancer in the patient.

As described above, the cancerous state of human cervical cells is correlated with changes in the levels of expression of the markers of the invention. The
5 invention includes a method for assessing the human cervical cell carcinogenic potential of a test compound. This method comprises maintaining separate aliquots of human cervical cells in the presence and absence of the test compound. Expression of a marker of the invention in each of the aliquots is compared. A significantly higher level of expression of a marker of the invention in the aliquot maintained in the presence of the
10 test compound (relative to the aliquot maintained in the absence of the test compound) is an indication that the test compound possesses human cervical cell carcinogenic potential. The relative carcinogenic potentials of various test compounds can be assessed by comparing the degree of enhancement or inhibition of the level of expression of the relevant markers, by comparing the number of markers for which the
15 level of expression is enhanced or inhibited, or by comparing both.

Various aspects of the invention are described in further detail in the following subsections.

I. Isolated Nucleic Acid Molecules
20 One aspect of the invention pertains to isolated nucleic acid molecules, including nucleic acids which encode a marker protein or a portion thereof. Isolated nucleic acids of the invention also include nucleic acid molecules sufficient for use as hybridization probes to identify marker nucleic acid molecules, and fragments of marker nucleic acid molecules, *e.g.*, those suitable for use as PCR primers for the amplification
25 or mutation of marker nucleic acid molecules. As used herein, the term "nucleic acid molecule" is intended to include DNA molecules (*e.g.*, cDNA or genomic DNA) and RNA molecules (*e.g.*, mRNA) and analogs of the DNA or RNA generated using nucleotide analogs. The nucleic acid molecule can be single-stranded or double-stranded, but preferably is double-stranded DNA.
30 An "isolated" nucleic acid molecule is one which is separated from other nucleic acid molecules which are present in the natural source of the nucleic acid molecule. Preferably, an "isolated" nucleic acid molecule is free of sequences (preferably protein-encoding sequences) which naturally flank the nucleic acid (*i.e.*,

sequences located at the 5' and 3' ends of the nucleic acid) in the genomic DNA of the organism from which the nucleic acid is derived. For example, in various embodiments, the isolated nucleic acid molecule can contain less than about 5 kB, 4 kB, 3 kB, 2 kB, 1 kB, 0.5 kB or 0.1 kB of nucleotide sequences which naturally flank the nucleic acid

5   molecule in genomic DNA of the cell from which the nucleic acid is derived. Moreover, an "isolated" nucleic acid molecule, such as a cDNA molecule, can be substantially free of other cellular material, or culture medium when produced by recombinant techniques, or substantially free of chemical precursors or other chemicals when chemically synthesized.

10   A nucleic acid molecule of the present invention can be isolated using standard molecular biology techniques and the sequence information in the database records described herein. Using all or a portion of such nucleic acid sequences, nucleic acid molecules of the invention can be isolated using standard hybridization and cloning techniques (*e.g.*, as described in Sambrook *et al.*, ed., *Molecular Cloning: A Laboratory*

15   *Manual, 2nd ed.,* Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989).

                A nucleic acid molecule of the invention can be amplified using cDNA, mRNA, or genomic DNA as a template and appropriate oligonucleotide primers according to standard PCR amplification techniques. The nucleic acid so amplified can

20   be cloned into an appropriate vector and characterized by DNA sequence analysis. Furthermore, nucleotides corresponding to all or a portion of a nucleic acid molecule of the invention can be prepared by standard synthetic techniques, *e.g.*, using an automated DNA synthesizer.

                In another preferred embodiment, an isolated nucleic acid molecule of the

25   invention comprises a nucleic acid molecule which has a nucleotide sequence complementary to the nucleotide sequence of a marker nucleic acid or to the nucleotide sequence of a nucleic acid encoding a marker protein. A nucleic acid molecule which is complementary to a given nucleotide sequence is one which is sufficiently complementary to the given nucleotide sequence that it can hybridize to the given

30   nucleotide sequence thereby forming a stable duplex.

                Moreover, a nucleic acid molecule of the invention can comprise only a portion of a nucleic acid sequence, wherein the full length nucleic acid sequence comprises a marker nucleic acid or which encodes a marker protein. Such nucleic acids

can be used, for example, as a probe or primer. The probe/primer typically is used as one or more substantially purified oligonucleotides. The oligonucleotide typically comprises a region of nucleotide sequence that hybridizes under stringent conditions to at least about 7, preferably about 15, more preferably about 25, 50, 75, 100, 125, 150,

5   175, 200, 250, 300, 350, or 400 or more consecutive nucleotides of a nucleic acid of the invention.

Probes based on the sequence of a nucleic acid molecule of the invention can be used to detect transcripts or genomic sequences corresponding to one or more markers of the invention. The probe comprises a label group attached thereto, e.g., a

10   radioisotope, a fluorescent compound, an enzyme, or an enzyme co-factor. Such probes can be used as part of a diagnostic test kit for identifying cells or tissues which mis-express the protein, such as by measuring levels of a nucleic acid molecule encoding the protein in a sample of cells from a subject, e.g., detecting mRNA levels or determining whether a gene encoding the protein has been mutated or deleted.

15   The invention further encompasses nucleic acid molecules that differ, due to degeneracy of the genetic code, from the nucleotide sequence of nucleic acids encoding a marker protein (e.g., a protein having one of the amino acid sequences set forth in the Sequence Listing), and thus encode the same protein.

It will be appreciated by those skilled in the art that DNA sequence

20   polymorphisms that lead to changes in the amino acid sequence can exist within a population (e.g., the human population). Such genetic polymorphisms can exist among individuals within a population due to natural allelic variation. An allele is one of a group of genes which occur alternatively at a given genetic locus. In addition, it will be appreciated that DNA polymorphisms that affect RNA expression levels can also exist

25   that may affect the overall expression level of that gene (e.g., by affecting regulation or degradation).

As used herein, the phrase "allelic variant" refers to a nucleotide sequence which occurs at a given locus or to a polypeptide encoded by the nucleotide sequence.

As used herein, the terms "gene" and "recombinant gene" refer to nucleic

30   acid molecules comprising an open reading frame encoding a polypeptide corresponding to a marker of the invention. Such natural allelic variations can typically result in 1-5% variance in the nucleotide sequence of a given gene. Alternative alleles can be identified by sequencing the gene of interest in a number of different individuals. This can be

readily carried out by using hybridization probes to identify the same genetic locus in a variety of individuals. Any and all such nucleotide variations and resulting amino acid polymorphisms or variations that are the result of natural allelic variation and that do not alter the functional activity are intended to be within the scope of the invention.

5   In another embodiment, an isolated nucleic acid molecule of the invention is at least 7, 15, 20, 25, 30, 40, 60, 80, 100, 150, 200, 250, 300, 350, 400, 450, 550, 650, 700, 800, 900, 1000, 1200, 1400, 1600, 1800, 2000, 2200, 2400, 2600, 2800, 3000, 3500, 4000, 4500, or more nucleotides in length and hybridizes under stringent conditions to a marker nucleic acid or to a nucleic acid encoding a marker protein. As

10   used herein, the term "hybridizes under stringent conditions" is intended to describe conditions for hybridization and washing under which nucleotide sequences at least 60% (65%, 70%, preferably 75%) identical to each other typically remain hybridized to each other. Such stringent conditions are known to those skilled in the art and can be found in sections 6.3.1-6.3.6 of *Current Protocols in Molecular Biology*, John Wiley & Sons,

15   N.Y. (1989). A preferred, non-limiting example of stringent hybridization conditions are hybridization in 6X sodium chloride/sodium citrate (SSC) at about 45°C, followed by one or more washes in 0.2X SSC, 0.1% SDS at 50-65°C.

          In addition to naturally-occurring allelic variants of a nucleic acid molecule of the invention that can exist in the population, the skilled artisan will further

20   appreciate that sequence changes can be introduced by mutation thereby leading to changes in the amino acid sequence of the encoded protein, without altering the biological activity of the protein encoded thereby. For example, one can make nucleotide substitutions leading to amino acid substitutions at "non-essential" amino acid residues. A "non-essential" amino acid residue is a residue that can be altered from

25   the wild-type sequence without altering the biological activity, whereas an "essential" amino acid residue is required for biological activity. For example, amino acid residues that are not conserved or only semi-conserved among homologs of various species may be non-essential for activity and thus would be likely targets for alteration. Alternatively, amino acid residues that are conserved among the homologs of various

30   species (*e.g.*, murine and human) may be essential for activity and thus would not be likely targets for alteration.

Accordingly, another aspect of the invention pertains to nucleic acid molecules encoding a variant marker protein that contain changes in amino acid residues that are not essential for activity. Such variant marker proteins differ in amino acid sequence from the naturally-occurring marker proteins, yet retain biological activity. In

5   one embodiment, such a variant marker protein has an amino acid sequence that is at least about 40% identical, 50%, 60%, 70%, 80%, 90%, 95%, or 98% identical to the amino acid sequence of a marker protein.

An isolated nucleic acid molecule encoding a variant marker protein can be created by introducing one or more nucleotide substitutions, additions or deletions

10   into the nucleotide sequence of marker nucleic acids, such that one or more amino acid residue substitutions, additions, or deletions are introduced into the encoded protein. Mutations can be introduced by standard techniques, such as site-directed mutagenesis and PCR-mediated mutagenesis. Preferably, conservative amino acid substitutions are made at one or more predicted non-essential amino acid residues. A "conservative

15   amino acid substitution" is one in which the amino acid residue is replaced with an amino acid residue having a similar side chain. Families of amino acid residues having similar side chains have been defined in the art. These families include amino acids with basic side chains (e.g., lysine, arginine, histidine), acidic side chains (e.g., aspartic acid, glutamic acid), uncharged polar side chains (e.g., glycine, asparagine, glutamine,

20   serine, threonine, tyrosine, cysteine), non-polar side chains (e.g., alanine, valine, leucine, isoleucine, proline, phenylalanine, methionine, tryptophan), beta-branched side chains (e.g., threonine, valine, isoleucine) and aromatic side chains (e.g., tyrosine, phenylalanine, tryptophan, histidine). Alternatively, mutations can be introduced randomly along all or part of the coding sequence, such as by saturation mutagenesis,

25   and the resultant mutants can be screened for biological activity to identify mutants that retain activity. Following mutagenesis, the encoded protein can be expressed recombinantly and the activity of the protein can be determined.

The present invention encompasses antisense nucleic acid molecules, i.e., molecules which are complementary to a sense nucleic acid of the invention, e.g.,

30   complementary to the coding strand of a double-stranded marker cDNA molecule or complementary to a marker mRNA sequence. Accordingly, an antisense nucleic acid of the invention can hydrogen bond to (i.e. anneal with) a sense nucleic acid of the invention. The antisense nucleic acid can be complementary to an entire coding strand,

or to only a portion thereof, *e.g.*, all or part of the protein coding region (or open reading frame). An antisense nucleic acid molecule can also be antisense to all or part of a non-coding region of the coding strand of a nucleotide sequence encoding a marker protein. The non-coding regions ("5' and 3' untranslated regions") are the 5' and 3' sequences

5   which flank the coding region and are not translated into amino acids.

An antisense oligonucleotide can be, for example, about 5, 10, 15, 20, 25, 30, 35, 40, 45, or 50 or more nucleotides in length. An antisense nucleic acid of the invention can be constructed using chemical synthesis and enzymatic ligation reactions using procedures known in the art. For example, an antisense nucleic acid (*e.g.*, an

10   antisense oligonucleotide) can be chemically synthesized using naturally occurring nucleotides or variously modified nucleotides designed to increase the biological stability of the molecules or to increase the physical stability of the duplex formed between the antisense and sense nucleic acids, *e.g.*, phosphorothioate derivatives and acridine substituted nucleotides can be used. Examples of modified nucleotides which

15   can be used to generate the antisense nucleic acid include 5-fluorouracil, 5-bromouracil, 5-chlorouracil, 5-iodouracil, hypoxanthine, xanthine, 4-acetylcytosine, 5-(carboxyhydroxylmethyl) uracil, 5-carboxymethylaminomethyl-2-thiouridine, 5-carboxymethylaminomethyluracil, dihydrouracil, beta-D-galactosylqueosine, inosine, N6-isopentenyladenine, 1-methylguanine, 1-methylinosine, 2,2-dimethylguanine, 2-

20   methyladenine, 2-methylguanine, 3-methylcytosine, 5-methylcytosine, N6-adenine, 7-methylguanine, 5-methylaminomethyluracil, 5-methoxyaminomethyl-2-thiouracil, beta-D-mannosylqueosine, 5'-methoxycarboxymethyluracil, 5-methoxyuracil, 2-methylthio-N6-isopentenyladenine, uracil-5-oxyacetic acid (v), wybutoxosine, pseudouracil, queosine, 2-thiocytosine, 5-methyl-2-thiouracil, 2-thiouracil, 4-thiouracil, 5-

25   methyluracil, uracil-5-oxyacetic acid methylester, uracil-5-oxyacetic acid (v), 5-methyl-2-thiouracil, 3-(3-amino-3-N-2-carboxypropyl) uracil, (acp3)w, and 2,6-diaminopurine. Alternatively, the antisense nucleic acid can be produced biologically using an expression vector into which a nucleic acid has been sub-cloned in an antisense orientation (*i.e.*, RNA transcribed from the inserted nucleic acid will be of an antisense

30   orientation to a target nucleic acid of interest, described further in the following subsection).

The antisense nucleic acid molecules of the invention are typically
administered to a subject or generated *in situ* such that they hybridize with or bind to
cellular mRNA and/or genomic DNA encoding a marker protein to thereby inhibit
expression of the marker, *e.g.*, by inhibiting transcription and/or translation. The
5   hybridization can be by conventional nucleotide complementarity to form a stable
duplex, or, for example, in the case of an antisense nucleic acid molecule which binds to
DNA duplexes, through specific interactions in the major groove of the double helix.
Examples of a route of administration of antisense nucleic acid molecules of the
invention includes direct injection at a tissue site or infusion of the antisense nucleic acid
10 into an ovary-associated body fluid. Alternatively, antisense nucleic acid molecules can
be modified to target selected cells and then administered systemically. For example,
for systemic administration, antisense molecules can be modified such that they
specifically bind to receptors or antigens expressed on a selected cell surface, *e.g.*, by
linking the antisense nucleic acid molecules to peptides or antibodies which bind to cell
15 surface receptors or antigens. The antisense nucleic acid molecules can also be
delivered to cells using the vectors described herein. To achieve sufficient intracellular
concentrations of the antisense molecules, vector constructs in which the antisense
nucleic acid molecule is placed under the control of a strong pol II or pol III promoter
are preferred.

20   An antisense nucleic acid molecule of the invention can be an α-anomeric
nucleic acid molecule. An α-anomeric nucleic acid molecule forms specific double-
stranded hybrids with complementary RNA in which, contrary to the usual α-units, the
strands run parallel to each other (Gaultier *et al.*, 1987, *Nucleic Acids Res.* 15:6625-
6641). The antisense nucleic acid molecule can also comprise a 2'-o-
25 methylribonucleotide (Inoue *et al.*, 1987, *Nucleic Acids Res.* 15:6131-6148) or a
chimeric RNA-DNA analogue (Inoue *et al.*, 1987, *FEBS Lett.* 215:327-330).

        The invention also encompasses ribozymes. Ribozymes are catalytic
RNA molecules with ribonuclease activity which are capable of cleaving a single-
stranded nucleic acid, such as an mRNA, to which they have a complementary region.
30 Thus, ribozymes (*e.g.*, hammerhead ribozymes as described in Haselhoff and Gerlach,
1988, *Nature* 334:585-591) can be used to catalytically cleave mRNA transcripts to
thereby inhibit translation of the protein encoded by the mRNA. A ribozyme having
specificity for a nucleic acid molecule encoding a marker protein can be designed based

upon the nucleotide sequence of a cDNA corresponding to the marker. For example, a derivative of a *Tetrahymena* L-19 IVS RNA can be constructed in which the nucleotide sequence of the active site is complementary to the nucleotide sequence to be cleaved (see Cech *et al.* U.S. Patent No. 4,987,071; and Cech *et al.* U.S. Patent No. 5,116,742).

5   Alternatively, an mRNA encoding a polypeptide of the invention can be used to select a catalytic RNA having a specific ribonuclease activity from a pool of RNA molecules (see, *e.g.*, Bartel and Szostak, 1993, *Science* 261:1411-1418).

The invention also encompasses nucleic acid molecules which form triple helical structures. For example, expression of a marker of the invention can be inhibited

10   by targeting nucleotide sequences complementary to the regulatory region of the gene encoding the marker nucleic acid or protein (*e.g.*, the promoter and/or enhancer) to form triple helical structures that prevent transcription of the gene in target cells. See generally Helene (1991) *Anticancer Drug Des.* 6(6):569-84; Helene (1992) *Ann. N.Y. Acad. Sci.* 660:27-36; and Maher (1992) *Bioassays* 14(12):807-15.

15   In various embodiments, the nucleic acid molecules of the invention can be modified at the base moiety, sugar moiety or phosphate backbone to improve, *e.g.*, the stability, hybridization, or solubility of the molecule. For example, the deoxyribose phosphate backbone of the nucleic acids can be modified to generate peptide nucleic acids (see Hyrup *et al.*, 1996, *Bioorganic & Medicinal Chemistry* 4(1): 5-23). As used

20   herein, the terms "peptide nucleic acids" or "PNAs" refer to nucleic acid mimics, *e.g.*, DNA mimics, in which the deoxyribose phosphate backbone is replaced by a pseudopeptide backbone and only the four natural nucleobases are retained. The neutral backbone of PNAs has been shown to allow for specific hybridization to DNA and RNA under conditions of low ionic strength. The synthesis of PNA oligomers can be

25   performed using standard solid phase peptide synthesis protocols as described in Hyrup *et al.* (1996), *supra*; Perry-O'Keefe *et al.* (1996) *Proc. Natl. Acad. Sci. USA* 93:14670-675.

PNAs can be used in therapeutic and diagnostic applications. For example, PNAs can be used as antisense or antigene agents for sequence-specific

30   modulation of gene expression by, *e.g.*, inducing transcription or translation arrest or inhibiting replication. PNAs can also be used, *e.g.*, in the analysis of single base pair mutations in a gene by, *e.g.*, PNA directed PCR clamping; as artificial restriction enzymes when used in combination with other enzymes, *e.g.*, S1 nucleases (Hyrup

(1996), *supra*; or as probes or primers for DNA sequence and hybridization (Hyrup, 1996, *supra*; Perry-O'Keefe *et al.*, 1996, *Proc. Natl. Acad. Sci. USA* 93:14670-675).

In another embodiment, PNAs can be modified, *e.g.*, to enhance their stability or cellular uptake, by attaching lipophilic or other helper groups to PNA, by the formation of PNA-DNA chimeras, or by the use of liposomes or other techniques of drug delivery known in the art. For example, PNA-DNA chimeras can be generated which can combine the advantageous properties of PNA and DNA. Such chimeras allow DNA recognition enzymes, *e.g.*, RNase H and DNA polymerases, to interact with the DNA portion while the PNA portion would provide high binding affinity and specificity. PNA-DNA chimeras can be linked using linkers of appropriate lengths selected in terms of base stacking, number of bonds between the nucleobases, and orientation (Hyrup, 1996, *supra*). The synthesis of PNA-DNA chimeras can be performed as described in Hyrup (1996), *supra*, and Finn *et al.* (1996) *Nucleic Acids Res.* 24(17):3357-63. For example, a DNA chain can be synthesized on a solid support using standard phosphoramidite coupling chemistry and modified nucleoside analogs. Compounds such as 5'-(4-methoxytrityl)amino-5'-deoxy-thymidine phosphoramidite can be used as a link between the PNA and the 5' end of DNA (Mag *et al.*, 1989, *Nucleic Acids Res.* 17:5973-88). PNA monomers are then coupled in a step-wise manner to produce a chimeric molecule with a 5' PNA segment and a 3' DNA segment (Finn *et al.*, 1996, *Nucleic Acids Res.* 24(17):3357-63). Alternatively, chimeric molecules can be synthesized with a 5' DNA segment and a 3' PNA segment (Peterser *et al.*, 1975, *Bioorganic Med. Chem. Lett.* 5:1119-11124).

In other embodiments, the oligonucleotide can include other appended groups such as peptides (*e.g.*, for targeting host cell receptors *in vivo*), or agents facilitating transport across the cell membrane (see, *e.g.*, Letsinger *et al.*, 1989, *Proc. Natl. Acad. Sci. USA* 86:6553-6556; Lemaitre *et al.*, 1987, *Proc. Natl. Acad. Sci. USA* 84:648-652; PCT Publication No. WO 88/09810) or the blood-brain barrier (see, *e.g.*, PCT Publication No. WO 89/10134). In addition, oligonucleotides can be modified with hybridization-triggered cleavage agents (see, *e.g.*, Krol *et al.*, 1988, *Bio/Techniques* 6:958-976) or intercalating agents (see, *e.g.*, Zon, 1988, *Pharm. Res.* 5:539-549). To this end, the oligonucleotide can be conjugated to another molecule, *e.g.*, a peptide, hybridization triggered cross-linking agent, transport agent, hybridization-triggered cleavage agent, etc.

The invention also includes molecular beacon nucleic acids having at least one region which is complementary to a nucleic acid of the invention, such that the molecular beacon is useful for quantitating the presence of the nucleic acid of the invention in a sample. A "molecular beacon" nucleic acid is a nucleic acid comprising a

5   pair of complementary regions and having a fluorophore and a fluorescent quencher associated therewith. The fluorophore and quencher are associated with different portions of the nucleic acid in such an orientation that when the complementary regions are annealed with one another, fluorescence of the fluorophore is quenched by the quencher. When the complementary regions of the nucleic acid are not annealed with

10   one another, fluorescence of the fluorophore is quenched to a lesser degree. Molecular beacon nucleic acids are described, for example, in U.S. Patent 5,876,930.

II. Isolated Proteins and Antibodies

One aspect of the invention pertains to isolated marker proteins and

15   biologically active portions thereof, as well as polypeptide fragments suitable for use as immunogens to raise antibodies directed against a marker protein or a fragment thereof. In one embodiment, the native marker protein can be isolated from cells or tissue sources by an appropriate purification scheme using standard protein purification techniques. In another embodiment, a protein or peptide comprising the whole or a

20   segment of the marker protein is produced by recombinant DNA techniques. Alternative to recombinant expression, such protein or peptide can be synthesized chemically using standard peptide synthesis techniques.

An "isolated" or "purified" protein or biologically active portion thereof is substantially free of cellular material or other contaminating proteins from the cell or

25   tissue source from which the protein is derived, or substantially free of chemical precursors or other chemicals when chemically synthesized. The language "substantially free of cellular material" includes preparations of protein in which the protein is separated from cellular components of the cells from which it is isolated or recombinantly produced. Thus, protein that is substantially free of cellular material

30   includes preparations of protein having less than about 30%, 20%, 10%, or 5% (by dry weight) of heterologous protein (also referred to herein as a "contaminating protein"). When the protein or biologically active portion thereof is recombinantly produced, it is also preferably substantially free of culture medium, i.e., culture medium represents less

than about 20%, 10%, or 5% of the volume of the protein preparation. When the protein
is produced by chemical synthesis, it is preferably substantially free of chemical
precursors or other chemicals, *i.e.*, it is separated from chemical precursors or other
chemicals which are involved in the synthesis of the protein. Accordingly such

5   preparations of the protein have less than about 30%, 20%, 10%, 5% (by dry weight) of
chemical precursors or compounds other than the polypeptide of interest.

Biologically active portions of a marker protein include polypeptides
comprising amino acid sequences sufficiently identical to or derived from the amino
acid sequence of the marker protein, which include fewer amino acids than the full

10   length protein, and exhibit at least one activity of the corresponding full-length protein.
Typically, biologically active portions comprise a domain or motif with at least one
activity of the corresponding full-length protein. A biologically active portion of a
marker protein of the invention can be a polypeptide which is, for example, 10, 25, 50,
100 or more amino acids in length. Moreover, other biologically active portions, in

15   which other regions of the marker protein are deleted, can be prepared by recombinant
techniques and evaluated for one or more of the functional activities of the native form
of the marker protein.

Preferred marker proteins are encoded by nucleotide sequences
comprising the sequence of any of the sequences set forth in the Sequence Listing.

20   Other useful proteins are substantially identical (*e.g.*, at least about 40%, preferably
50%, 60%, 70%, 80%, 90%, 95%, or 99%) to one of these sequences and retain the
functional activity of the corresponding naturally-occurring marker protein yet differ in
amino acid sequence due to natural allelic variation or mutagenesis.

To determine the percent identity of two amino acid sequences or of two

25   nucleic acids, the sequences are aligned for optimal comparison purposes (*e.g.*, gaps can
be introduced in the sequence of a first amino acid or nucleic acid sequence for optimal
alignment with a second amino or nucleic acid sequence). The amino acid residues or
nucleotides at corresponding amino acid positions or nucleotide positions are then
compared. When a position in the first sequence is occupied by the same amino acid

30   residue or nucleotide as the corresponding position in the second sequence, then the
molecules are identical at that position. The percent identity between the two sequences
is a function of the number of identical positions shared by the sequences (*i.e.*, %

identity = # of identical positions/total # of positions (*e.g.*, overlapping positions) x100).
In one embodiment the two sequences are the same length.

The determination of percent identity between two sequences can be
accomplished using a mathematical algorithm. A preferred, non-limiting example of a

5   mathematical algorithm utilized for the comparison of two sequences is the algorithm of
Karlin and Altschul (1990) *Proc. Natl. Acad. Sci. USA* 87:2264-2268, modified as in
Karlin and Altschul (1993) *Proc. Natl. Acad. Sci. USA* 90:5873-5877. Such an
algorithm is incorporated into the BLASTN and BLASTX programs of Altschul, *et al.*
(1990) *J. Mol. Biol.* 215:403-410. BLAST nucleotide searches can be performed with

10   the BLASTN program, score = 100, wordlength = 12 to obtain nucleotide sequences
homologous to a nucleic acid molecules of the invention. BLAST protein searches can
be performed with the BLASTP program, score = 50, wordlength = 3 to obtain amino
acid sequences homologous to a protein molecules of the invention. To obtain gapped
alignments for comparison purposes, a newer version of the BLAST algorithm called

15   Gapped BLAST can be utilized as described in Altschul *et al.* (1997) *Nucleic Acids Res.*
25:3389-3402, which is able to perform gapped local alignments for the programs
BLASTN, BLASTP and BLASTX. Alternatively, PSI-Blast can be used to perform an
iterated search which detects distant relationships between molecules. When utilizing
BLAST, Gapped BLAST, and PSI-Blast programs, the default parameters of the

20   respective programs (*e.g.*, BLASTX and BLASTN) can be used. See
http://www.ncbi.nlm.nih.gov. Another preferred, non-limiting example of a
mathematical algorithm utilized for the comparison of sequences is the algorithm of
Myers and Miller, (1988) *CABIOS* 4:11-17. Such an algorithm is incorporated into the
ALIGN program (version 2.0) which is part of the GCG sequence alignment software

25   package. When utilizing the ALIGN program for comparing amino acid sequences, a
PAM120 weight residue table, a gap length penalty of 12, and a gap penalty of 4 can be
used. Yet another useful algorithm for identifying regions of local sequence similarity
and alignment is the FASTA algorithm as described in Pearson and Lipman (1988)
Proc. Natl. Acad. Sci. USA 85:2444-2448. When using the FASTA algorithm for

30   comparing nucleotide or amino acid sequences, a PAM120 weight residue table can, for
example, be used with a *k*-tuple value of 2.

The percent identity between two sequences can be determined using techniques similar to those described above, with or without allowing gaps. In calculating percent identity, only exact matches are counted.

The invention also provides chimeric or fusion proteins comprising a
5   marker protein or a segment thereof. As used herein, a "chimeric protein" or "fusion protein" comprises all or part (preferably a biologically active part) of a marker protein operably linked to a heterologous polypeptide (*i.e.*, a polypeptide other than the marker protein). Within the fusion protein, the term "operably linked" is intended to indicate that the marker protein or segment thereof and the heterologous polypeptide are fused
10   in-frame to each other. The heterologous polypeptide can be fused to the amino-terminus or the carboxyl-terminus of the marker protein or segment.

One useful fusion protein is a GST fusion protein in which a marker protein or segment is fused to the carboxyl terminus of GST sequences. Such fusion proteins can facilitate the purification of a recombinant polypeptide of the invention.
15   In another embodiment, the fusion protein contains a heterologous signal sequence at its amino terminus. For example, the native signal sequence of a marker protein can be removed and replaced with a signal sequence from another protein. For example, the gp67 secretory sequence of the baculovirus envelope protein can be used as a heterologous signal sequence (Ausubel *et al.*, ed., *Current Protocols in Molecular*
20   *Biology*, John Wiley & Sons, NY, 1992). Other examples of eukaryotic heterologous signal sequences include the secretory sequences of melittin and human placental alkaline phosphatase (Stratagene; La Jolla, California). In yet another example, useful prokaryotic heterologous signal sequences include the phoA secretory signal (Sambrook *et al.*, *supra*) and the protein A secretory signal (Pharmacia Biotech; Piscataway, New
25   Jersey).

In yet another embodiment, the fusion protein is an immunoglobulin fusion protein in which all or part of a marker protein is fused to sequences derived from a member of the immunoglobulin protein family. The immunoglobulin fusion proteins of the invention can be incorporated into pharmaceutical compositions and administered
30   to a subject to inhibit an interaction between a ligand (soluble or membrane-bound) and a protein on the surface of a cell (receptor), to thereby suppress signal transduction *in vivo*. The immunoglobulin fusion protein can be used to affect the bioavailability of a cognate ligand of a marker protein. Inhibition of ligand/receptor interaction can be

useful therapeutically, both for treating proliferative and differentiative disorders and for modulating (*e.g.* promoting or inhibiting) cell survival. Moreover, the immunoglobulin fusion proteins of the invention can be used as immunogens to produce antibodies directed against a marker protein in a subject, to purify ligands and in screening assays

5   to identify molecules which inhibit the interaction of the marker protein with ligands.

Chimeric and fusion proteins of the invention can be produced by standard recombinant DNA techniques. In another embodiment, the fusion gene can be synthesized by conventional techniques including automated DNA synthesizers. Alternatively, PCR amplification of gene fragments can be carried out using anchor

10   primers which give rise to complementary overhangs between two consecutive gene fragments which can subsequently be annealed and re-amplified to generate a chimeric gene sequence (see, *e.g.*, Ausubel *et al., supra*). Moreover, many expression vectors are commercially available that already encode a fusion moiety (*e.g.*, a GST polypeptide). A nucleic acid encoding a polypeptide of the invention can be cloned into such an

15   expression vector such that the fusion moiety is linked in-frame to the polypeptide of the invention.

A signal sequence can be used to facilitate secretion and isolation of marker proteins. Signal sequences are typically characterized by a core of hydrophobic amino acids which are generally cleaved from the mature protein during secretion in one

20   or more cleavage events. Such signal peptides contain processing sites that allow cleavage of the signal sequence from the mature proteins as they pass through the secretory pathway. Thus, the invention pertains to marker proteins, fusion proteins or segments thereof having a signal sequence, as well as to such proteins from which the signal sequence has been proteolytically cleaved (*i.e.*, the cleavage products). In one

25   embodiment, a nucleic acid sequence encoding a signal sequence can be operably linked in an expression vector to a protein of interest, such as a marker protein or a segment thereof. The signal sequence directs secretion of the protein, such as from a eukaryotic host into which the expression vector is transformed, and the signal sequence is subsequently or concurrently cleaved. The protein can then be readily purified from the

30   extracellular medium by art recognized methods. Alternatively, the signal sequence can be linked to the protein of interest using a sequence which facilitates purification, such as with a GST domain.

The present invention also pertains to variants of the marker proteins. Such variants have an altered amino acid sequence which can function as either agonists (mimetics) or as antagonists. Variants can be generated by mutagenesis, *e.g.*, discrete point mutation or truncation. An agonist can retain substantially the same, or a subset,

5   of the biological activities of the naturally occurring form of the protein. An antagonist of a protein can inhibit one or more of the activities of the naturally occurring form of the protein by, for example, competitively binding to a downstream or upstream member of a cellular signaling cascade which includes the protein of interest. Thus, specific biological effects can be elicited by treatment with a variant of limited function.

10   Treatment of a subject with a variant having a subset of the biological activities of the naturally occurring form of the protein can have fewer side effects in a subject relative to treatment with the naturally occurring form of the protein.

Variants of a marker protein which function as either agonists (mimetics) or as antagonists can be identified by screening combinatorial libraries of mutants, *e.g.*,

15   truncation mutants, of the protein of the invention for agonist or antagonist activity. In one embodiment, a variegated library of variants is generated by combinatorial mutagenesis at the nucleic acid level and is encoded by a variegated gene library. A variegated library of variants can be produced by, for example, enzymatically ligating a mixture of synthetic oligonucleotides into gene sequences such that a degenerate set of

20   potential protein sequences is expressible as individual polypeptides, or alternatively, as a set of larger fusion proteins (*e.g.*, for phage display). There are a variety of methods which can be used to produce libraries of potential variants of the marker proteins from a degenerate oligonucleotide sequence. Methods for synthesizing degenerate oligonucleotides are known in the art (see, *e.g.*, Narang, 1983, *Tetrahedron* 39:3; Itakura

25   *et al.*, 1984, *Annu. Rev. Biochem.* 53:323; Itakura *et al.*, 1984, *Science* 198:1056; Ike *et al.*, 1983 *Nucleic Acid Res.* 11:477).

In addition, libraries of segments of a marker protein can be used to generate a variegated population of polypeptides for screening and subsequent selection of variant marker proteins or segments thereof. For example, a library of coding

30   sequence fragments can be generated by treating a double stranded PCR fragment of the coding sequence of interest with a nuclease under conditions wherein nicking occurs only about once per molecule, denaturing the double stranded DNA, renaturing the DNA to form double stranded DNA which can include sense/antisense pairs from different

nicked products, removing single stranded portions from reformed duplexes by treatment with S1 nuclease, and ligating the resulting fragment library into an expression vector. By this method, an expression library can be derived which encodes amino terminal and internal fragments of various sizes of the protein of interest.

5   Several techniques are known in the art for screening gene products of combinatorial libraries made by point mutations or truncation, and for screening cDNA libraries for gene products having a selected property. The most widely used techniques, which are amenable to high through-put analysis, for screening large gene libraries typically include cloning the gene library into replicable expression vectors,

10   transforming appropriate cells with the resulting library of vectors, and expressing the combinatorial genes under conditions in which detection of a desired activity facilitates isolation of the vector encoding the gene whose product was detected. Recursive ensemble mutagenesis (REM), a technique which enhances the frequency of functional mutants in the libraries, can be used in combination with the screening assays to identify

15   variants of a protein of the invention (Arkin and Yourvan, 1992, *Proc. Natl. Acad. Sci. USA* 89:7811-7815; Delgrave *et al.,* 1993, *Protein Engineering* 6(3):327-331).

           Another aspect of the invention pertains to antibodies directed against a protein of the invention. In preferred embodiments, the antibodies specifically bind a marker protein or a fragment thereof. The terms "antibody" and "antibodies" as used

20   interchangeably herein refer to immunoglobulin molecules as well as fragments and derivatives thereof that comprise an immunologically active portion of an immunoglobulin molecule, (*i.e.,* such a portion contains an antigen binding site which specifically binds an antigen, such as a marker protein, *e.g.,* an epitope of a marker protein). An antibody which specifically binds to a protein of the invention is an

25   antibody which binds the protein, but does not substantially bind other molecules in a sample, *e.g.,* a biological sample, which naturally contains the protein. Examples of an immunologically active portion of an immunoglobulin molecule include, but are not limited to, single-chain antibodies (scAb), F(ab) and F(ab')$_2$ fragments.

           An isolated protein of the invention or a fragment thereof can be used as

30   an immunogen to generate antibodies. The full-length protein can be used or, alternatively, the invention provides antigenic peptide fragments for use as immunogens. The antigenic peptide of a protein of the invention comprises at least 8 (preferably 10, 15, 20, or 30 or more) amino acid residues of the amino acid sequence of one of the

proteins of the invention, and encompasses at least one epitope of the protein such that an antibody raised against the peptide forms a specific immune complex with the protein. Preferred epitopes encompassed by the antigenic peptide are regions that are located on the surface of the protein, *e.g.*, hydrophilic regions. Hydrophobicity sequence

5   analysis, hydrophilicity sequence analysis, or similar analyses can be used to identify hydrophilic regions. In preferred embodiments, an isolated marker protein or fragment thereof is used as an immunogen.

An immunogen typically is used to prepare antibodies by immunizing a suitable (*i.e.* immunocompetent) subject such as a rabbit, goat, mouse, or other mammal

10   or vertebrate. An appropriate immunogenic preparation can contain, for example, recombinantly-expressed or chemically-synthesized protein or peptide. The preparation can further include an adjuvant, such as Freund's complete or incomplete adjuvant, or a similar immunostimulatory agent. Preferred immunogen compositions are those that contain no other human proteins such as, for example, immunogen compositions made

15   using a non-human host cell for recombinant expression of a protein of the invention. In such a manner, the resulting antibody compositions have reduced or no binding of human proteins other than a protein of the invention.

The invention provides polyclonal and monoclonal antibodies. The term "monoclonal antibody" or "monoclonal antibody composition", as used herein, refers to

20   a population of antibody molecules that contain only one species of an antigen binding site capable of immunoreacting with a particular epitope. Preferred polyclonal and monoclonal antibody compositions are ones that have been selected for antibodies directed against a protein of the invention. Particularly preferred polyclonal and monoclonal antibody preparations are ones that contain only antibodies directed against

25   a marker protein or fragment thereof.

Polyclonal antibodies can be prepared by immunizing a suitable subject with a protein of the invention as an immunogen The antibody titer in the immunized subject can be monitored over time by standard techniques, such as with an enzyme linked immunosorbent assay (ELISA) using immobilized polypeptide. At an appropriate

30   time after immunization, *e.g.*, when the specific antibody titers are highest, antibody-producing cells can be obtained from the subject and used to prepare monoclonal antibodies (mAb) by standard techniques, such as the hybridoma technique originally described by Kohler and Milstein (1975) *Nature* 256:495-497, the human B cell

hybridoma technique (see Kozbor *et al.*, 1983, *Immunol. Today* 4:72), the EBV-hybridoma technique (see Cole *et al.*, pp. 77-96 In *Monoclonal Antibodies and Cancer Therapy*, Alan R. Liss, Inc., 1985) or trioma techniques. The technology for producing hybridomas is well known (see generally *Current Protocols in Immunology*, Coligan *et al.* ed., John Wiley & Sons, New York, 1994). Hybridoma cells producing a monoclonal antibody of the invention are detected by screening the hybridoma culture supernatants for antibodies that bind the polypeptide of interest, *e.g.*, using a standard ELISA assay.

Alternative to preparing monoclonal antibody-secreting hybridomas, a monoclonal antibody directed against a protein of the invention can be identified and isolated by screening a recombinant combinatorial immunoglobulin library (*e.g.*, an antibody phage display library) with the polypeptide of interest. Kits for generating and screening phage display libraries are commercially available (*e.g.*, the Pharmacia *Recombinant Phage Antibody System*, Catalog No. 27-9400-01; and the Stratagene *SurfZAP Phage Display Kit*, Catalog No. 240612). Additionally, examples of methods and reagents particularly amenable for use in generating and screening antibody display library can be found in, for example, U.S. Patent No. 5,223,409; PCT Publication No. WO 92/18619; PCT Publication No. WO 91/17271; PCT Publication No. WO 92/20791; PCT Publication No. WO 92/15679; PCT Publication No. WO 93/01288; PCT Publication No. WO 92/01047; PCT Publication No. WO 92/09690; PCT Publication No. WO 90/02809; Fuchs *et al.* (1991) *Bio/Technology* 9:1370-1372; Hay *et al.* (1992) *Hum. Antibod. Hybridomas* 3:81-85; Huse *et al.* (1989) *Science* 246:1275- 1281; Griffiths *et al.* (1993) *EMBO J.* 12:725-734.

The invention also provides recombinant antibodies that specifically bind a protein of the invention. In preferred embodiments, the recombinant antibodies specifically binds a marker protein or fragment thereof. Recombinant antibodies include, but are not limited to, chimeric and humanized monoclonal antibodies, comprising both human and non-human portions, single-chain antibodies and multi-specific antibodies. A chimeric antibody is a molecule in which different portions are derived from different animal species, such as those having a variable region derived from a murine mAb and a human immunoglobulin constant region. (See, *e.g.*, Cabilly et al., U.S. Patent No. 4,816,567; and Boss et al., U.S. Patent No. 4,816,397, which are incorporated herein by reference in their entirety.) Single-chain antibodies have an

antigen binding site and consist of a single polypeptide. They can be produced by techniques known in the art, for example using methods described in Ladner *et. al* U.S. Pat. No. 4,946,778 (which is incorporated herein by reference in its entirety); Bird *et al.*, (1988) *Science* 242:423-426; Whitlow *et al.*, (1991) *Methods in Enzymology* 2:1-9;

5   Whitlow *et al.*, (1991) *Methods in Enzymology* 2:97-105; and Huston *et al.*, (1991) *Methods in Enzymology Molecular Design and Modeling: Concepts and Applications* 203:46-88. Multi-specific antibodies are antibody molecules having at least two antigen-binding sites that specifically bind different antigens. Such molecules can be produced by techniques known in the art, for example using methods described in Segal,

10   U.S. Patent No. 4,676,980 (the disclosure of which is incorporated herein by reference in its entirety); Holliger et al., (1993) *Proc. Natl. Acad. Sci. USA* 90:6444-6448; *Whitlow et al.*, (1994) *Protein Eng.* 7:1017-1026 and U.S. Pat. No. 6,121,424.

Humanized antibodies are antibody molecules from non-human species having one or more complementarity determining regions (CDRs) from the non-human

15   species and a framework region from a human immunoglobulin molecule. (See, *e.g.*, Queen, U.S. Patent No. 5,585,089, which is incorporated herein by reference in its entirety.) Humanized monoclonal antibodies can be produced by recombinant DNA techniques known in the art, for example using methods described in PCT Publication No. WO 87/02671; European Patent Application 184,187; European Patent Application

20   171,496; European Patent Application 173,494; PCT Publication No. WO 86/01533; U.S. Patent No. 4,816,567; European Patent Application 125,023; Better *et al.* (1988) *Science* 240:1041-1043; Liu *et al.* (1987) *Proc. Natl. Acad. Sci. USA* 84:3439-3443; Liu *et al.* (1987) *J. Immunol.* 139:3521-3526; Sun *et al.* (1987) *Proc. Natl. Acad. Sci. USA* 84:214-218; Nishimura *et al.* (1987) *Cancer Res.* 47:999-1005; Wood *et al.* (1985)

25   *Nature* 314:446-449; and Shaw *et al.* (1988) *J. Natl. Cancer Inst.* 80:1553-1559); Morrison (1985) *Science* 229:1202-1207; Oi *et al.* (1986) *Bio/Techniques* 4:214; U.S. Patent 5,225,539; Jones *et al.* (1986) *Nature* 321:552-525; Verhoeyan *et al.* (1988) *Science* 239:1534; and Beidler *et al.* (1988) *J. Immunol.* 141:4053-4060.

More particularly, humanized antibodies can be produced, for example,

30   using transgenic mice which are incapable of expressing endogenous immunoglobulin heavy and light chains genes, but which can express human heavy and light chain genes. The transgenic mice are immunized in the normal fashion with a selected antigen, *e.g.*, all or a portion of a polypeptide corresponding to a marker of the invention. Monoclonal

antibodies directed against the antigen can be obtained using conventional hybridoma technology. The human immunoglobulin transgenes harbored by the transgenic mice rearrange during B cell differentiation, and subsequently undergo class switching and somatic mutation. Thus, using such a technique, it is possible to produce therapeutically

5   useful IgG, IgA and IgE antibodies. For an overview of this technology for producing human antibodies, see Lonberg and Huszar (1995) *Int. Rev. Immunol.* 13:65-93). For a detailed discussion of this technology for producing human antibodies and human monoclonal antibodies and protocols for producing such antibodies, see, *e.g.*, U.S. Patent 5,625,126; U.S. Patent 5,633,425; U.S. Patent 5,569,825; U.S. Patent 5,661,016;

10 and U.S. Patent 5,545,806. In addition, companies such as Abgenix, Inc. (Freemont, CA), can be engaged to provide human antibodies directed against a selected antigen using technology similar to that described above.

Completely human antibodies which recognize a selected epitope can be generated using a technique referred to as "guided selection." In this approach a selected

15 non-human monoclonal antibody, *e.g.*, a murine antibody, is used to guide the selection of a completely human antibody recognizing the same epitope (Jespers *et al.*, 1994, *Bio/technology* 12:899-903).

The antibodies of the invention can be isolated after production (*e.g.*, from the blood or serum of the subject) or synthesis and further purified by well-known

20 techniques. For example, IgG antibodies can be purified using protein A chromatography. Antibodies specific for a protein of the invention can be selected or (*e.g.*, partially purified) or purified by, *e.g.*, affinity chromatography. For example, a recombinantly expressed and purified (or partially purified) protein of the invention is produced as described herein, and covalently or non-covalently coupled to a solid

25 support such as, for example, a chromatography column. The column can then be used to affinity purify antibodies specific for the proteins of the invention from a sample containing antibodies directed against a large number of different epitopes, thereby generating a substantially purified antibody composition, *i.e.*, one that is substantially free of contaminating antibodies. By a substantially purified antibody composition is

30 meant, in this context, that the antibody sample contains at most only 30% (by dry weight) of contaminating antibodies directed against epitopes other than those of the desired protein of the invention, and preferably at most 20%, yet more preferably at most 10%, and most preferably at most 5% (by dry weight) of the sample is

contaminating antibodies. A purified antibody composition means that at least 99% of the antibodies in the composition are directed against the desired protein of the invention.

In a preferred embodiment, the substantially purified antibodies of the invention may specifically bind to a signal peptide, a secreted sequence, an extracellular domain, a transmembrane or a cytoplasmic domain or cytoplasmic membrane of a protein of the invention. In a particularly preferred embodiment, the substantially purified antibodies of the invention specifically bind to a secreted sequence or an extracellular domain of the amino acid sequences of a protein of the invention. In a more preferred embodiment, the substantially purified antibodies of the invention specifically bind to a secreted sequence or an extracellular domain of the amino acid sequences of a marker protein.

An antibody directed against a protein of the invention can be used to isolate the protein by standard techniques, such as affinity chromatography or immunoprecipitation. Moreover, such an antibody can be used to detect the marker protein or fragment thereof (*e.g.*, in a cellular lysate or cell supernatant) in order to evaluate the level and pattern of expression of the marker. The antibodies can also be used diagnostically to monitor protein levels in tissues or body fluids (*e.g.* in a cervical-associated body fluid) as part of a clinical testing procedure, *e.g.*, to, for example, determine the efficacy of a given treatment regimen. Detection can be facilitated by the use of an antibody derivative, which comprises an antibody of the invention coupled to a detectable substance. Examples of detectable substances include various enzymes, prosthetic groups, fluorescent materials, luminescent materials, bioluminescent materials, and radioactive materials. Examples of suitable enzymes include horseradish peroxidase, alkaline phosphatase, β-galactosidase, or acetylcholinesterase; examples of suitable prosthetic group complexes include streptavidin/biotin and avidin/biotin; examples of suitable fluorescent materials include umbelliferone, fluorescein, fluorescein isothiocyanate, rhodamine, dichlorotriazinylamine fluorescein, dansyl chloride or phycoerythrin; an example of a luminescent material includes luminol; examples of bioluminescent materials include luciferase, luciferin, and aequorin, and examples of suitable radioactive material include ^{125}I, ^{131}I, ^{35}S or ^{3}H.

Antibodies of the invention may also be used as therapeutic agents in treating cancers. In a preferred embodiment, completely human antibodies of the invention are used for therapeutic treatment of human cancer patients, particularly those having an cervical cancer. In another preferred embodiment, antibodies that bind

5   specifically to a marker protein or fragment thereof are used for therapeutic treatment. Further, such therapeutic antibody may be an antibody derivative or immunotoxin comprising an antibody conjugated to a therapeutic moiety such as a cytotoxin, a therapeutic agent or a radioactive metal ion. A cytotoxin or cytotoxic agent includes any agent that is detrimental to cells. Examples include taxol, cytochalasin B, gramicidin D,

10   ethidium bromide, emetine, mitomycin, etoposide, tenoposide, vincristine, vinblastine, colchicin, doxorubicin, daunorubicin, dihydroxy anthracin dione, mitoxantrone, mithramycin, actinomycin D, 1-dehydrotestosterone, glucocorticoids, procaine, tetracaine, lidocaine, propranolol, and puromycin and analogs or homologs thereof. Therapeutic agents include, but are not limited to, antimetabolites (e.g., methotrexate,

15   6-mercaptopurine, 6-thioguanine, cytarabine, 5-fluorouracil decarbazine), alkylating agents (e.g., mechlorethamine, thioepa chlorambucil, melphalan, carmustine (BSNU) and lomustine (CCNU), cyclothosphamide, busulfan, dibromomannitol, streptozotocin, mitomycin C, and cis-dichlorodiamine platinum (II) (DDP) cisplatin), anthracyclines (e.g., daunorubicin (formerly daunomycin) and doxorubicin), antibiotics (e.g.,

20   dactinomycin (formerly actinomycin), bleomycin, mithramycin, and anthramycin (AMC)), and anti-mitotic agents (e.g., vincristine and vinblastine).

The conjugated antibodies of the invention can be used for modifying a given biological response, for the drug moiety is not to be construed as limited to classical chemical therapeutic agents. For example, the drug moiety may be a protein or

25   polypeptide possessing a desired biological activity. Such proteins may include, for example, a toxin such as ribosome-inhibiting protein (see Better et al., U.S. Patent No. 6,146,631, the disclosure of which is incorporated herein in its entirety), abrin, ricin A, pseudomonas exotoxin, or diphtheria toxin; a protein such as tumor necrosis factor, .alpha.-interferon, .beta.-interferon, nerve growth factor, platelet derived growth factor,

30   tissue plasminogen activator; or, biological response modifiers such as, for example, lymphokines, interleukin-1 ("IL-1"), interleukin-2 ("IL-2"), interleukin-6 ("IL-6"), granulocyte macrophase colony stimulating factor ("GM-CSF"), granulocyte colony stimulating factor ("G-CSF"), or other growth factors.

Techniques for conjugating such therapeutic moiety to antibodies are well known, see, *e.g.*, Arnon et al., "Monoclonal Antibodies For Immunotargeting Of Drugs In Cancer Therapy", in Monoclonal Antibodies And Cancer Therapy, Reisfeld et al. (eds.), pp. 243-56 (Alan R. Liss, Inc. 1985); Hellstrom et al., "Antibodies For Drug Delivery", in Controlled Drug Delivery (2nd Ed.), Robinson et al. (eds.), pp. 623-53 (Marcel Dekker, Inc. 1987); Thorpe, "Antibody Carriers Of Cytotoxic Agents In Cancer Therapy: A Review", in Monoclonal Antibodies '84: Biological And Clinical Applications, Pinchera et al. (eds.), pp. 475-506 (1985); "Analysis, Results, And Future Prospective Of The Therapeutic Use Of Radiolabeled Antibody In Cancer Therapy", in Monoclonal Antibodies For Cancer Detection And Therapy, Baldwin et al. (eds.), pp. 303-16 (Academic Press 1985), and Thorpe et al., "The Preparation And Cytotoxic Properties Of Antibody-Toxin Conjugates", Immunol. Rev., 62:119-58 (1982).

Accordingly, in one aspect, the invention provides substantially purified antibodies, antibody fragments and derivatives, all of which specifically bind to a protein of the invention and preferably, a marker protein. In various embodiments, the substantially purified antibodies of the invention, or fragments or derivatives thereof, can be human, non-human, chimeric and/or humanized antibodies. In another aspect, the invention provides non-human antibodies, antibody fragments and derivatives, all of which specifically bind to a protein of the invention and preferably, a marker protein. Such non-human antibodies can be goat, mouse, sheep, horse, chicken, rabbit, or rat antibodies. Alternatively, the non-human antibodies of the invention can be chimeric and/or humanized antibodies. In addition, the non-human antibodies of the invention can be polyclonal antibodies or monoclonal antibodies. In still a further aspect, the invention provides monoclonal antibodies, antibody fragments and derivatives, all of which specifically bind to a protein of the invention and preferably, a marker protein. The monoclonal antibodies can be human, humanized, chimeric and/or non-human antibodies.

The invention also provides a kit containing an antibody of the invention conjugated to a detectable substance, and instructions for use. Still another aspect of the invention is a pharmaceutical composition comprising an antibody of the invention. In one embodiment, the pharmaceutical composition comprises an antibody of the invention and a pharmaceutically acceptable carrier.

III. Recombinant Expression Vectors and Host Cells

Another aspect of the invention pertains to vectors, preferably expression vectors, containing a nucleic acid encoding a marker protein (or a portion of such a protein). As used herein, the term "vector" refers to a nucleic acid molecule capable of transporting another nucleic acid to which it has been linked. One type of vector is a "plasmid", which refers to a circular double stranded DNA loop into which additional DNA segments can be ligated. Another type of vector is a viral vector, wherein additional DNA segments can be ligated into the viral genome. Certain vectors are capable of autonomous replication in a host cell into which they are introduced (e.g., bacterial vectors having a bacterial origin of replication and episomal mammalian vectors). Other vectors (e.g., non-episomal mammalian vectors) are integrated into the genome of a host cell upon introduction into the host cell, and thereby are replicated along with the host genome. Moreover, certain vectors, namely expression vectors, are capable of directing the expression of genes to which they are operably linked. In general, expression vectors of utility in recombinant DNA techniques are often in the form of plasmids (vectors). However, the invention is intended to include such other forms of expression vectors, such as viral vectors (e.g., replication defective retroviruses, adenoviruses and adeno-associated viruses), which serve equivalent functions.

The recombinant expression vectors of the invention comprise a nucleic acid of the invention in a form suitable for expression of the nucleic acid in a host cell. This means that the recombinant expression vectors include one or more regulatory sequences, selected on the basis of the host cells to be used for expression, which is operably linked to the nucleic acid sequence to be expressed. Within a recombinant expression vector, "operably linked" is intended to mean that the nucleotide sequence of interest is linked to the regulatory sequence(s) in a manner which allows for expression of the nucleotide sequence (e.g., in an in vitro transcription/translation system or in a host cell when the vector is introduced into the host cell). The term "regulatory sequence" is intended to include promoters, enhancers and other expression control elements (e.g., polyadenylation signals). Such regulatory sequences are described, for example, in Goeddel, Methods in Enzymology: Gene Expression Technology vol.185, Academic Press, San Diego, CA (1991). Regulatory sequences include those which direct constitutive expression of a nucleotide sequence in many types of host cell and

those which direct expression of the nucleotide sequence only in certain host cells (*e.g.*, tissue-specific regulatory sequences). It will be appreciated by those skilled in the art that the design of the expression vector can depend on such factors as the choice of the host cell to be transformed, the level of expression of protein desired, and the like. The

5   expression vectors of the invention can be introduced into host cells to thereby produce proteins or peptides, including fusion proteins or peptides, encoded by nucleic acids as described herein.

The recombinant expression vectors of the invention can be designed for expression of a marker protein or a segment thereof in prokaryotic (*e.g.*, *E. coli*) or

10   eukaryotic cells (*e.g.*, insect cells {using baculovirus expression vectors}, yeast cells or mammalian cells). Suitable host cells are discussed further in Goeddel, *supra*. Alternatively, the recombinant expression vector can be transcribed and translated *in vitro*, for example using T7 promoter regulatory sequences and T7 polymerase.

Expression of proteins in prokaryotes is most often carried out in *E. coli*

15   with vectors containing constitutive or inducible promoters directing the expression of either fusion or non-fusion proteins. Fusion vectors add a number of amino acids to a protein encoded therein, usually to the amino terminus of the recombinant protein. Such fusion vectors typically serve three purposes: 1) to increase expression of recombinant protein; 2) to increase the solubility of the recombinant protein; and 3) to aid in the

20   purification of the recombinant protein by acting as a ligand in affinity purification. Often, in fusion expression vectors, a proteolytic cleavage site is introduced at the junction of the fusion moiety and the recombinant protein to enable separation of the recombinant protein from the fusion moiety subsequent to purification of the fusion protein. Such enzymes, and their cognate recognition sequences, include Factor Xa,

25   thrombin and enterokinase. Typical fusion expression vectors include pGEX (Pharmacia Biotech Inc; Smith and Johnson, 1988, *Gene* 67:31-40), pMAL (New England Biolabs, Beverly, MA) and pRIT5 (Pharmacia, Piscataway, NJ) which fuse glutathione S-transferase (GST), maltose E binding protein, or protein A, respectively, to the target recombinant protein.

30   Examples of suitable inducible non-fusion *E. coli* expression vectors include pTrc (Amann *et al.*, 1988, *Gene* 69:301-315) and pET 11d (Studier *et al.*, p. 60-89, In *Gene Expression Technology: Methods in Enzymology* vol.185, Academic Press, San Diego, CA, 1991). Target gene expression from the pTrc vector relies on host RNA

polymerase transcription from a hybrid trp-lac fusion promoter. Target gene expression from the pET 11d vector relies on transcription from a T7 gn10-lac fusion promoter mediated by a co-expressed viral RNA polymerase (T7 gn1). This viral polymerase is supplied by host strains BL21(DE3) or HMS174(DE3) from a resident prophage

5 harboring a T7 gn1 gene under the transcriptional control of the lacUV 5 promoter.

One strategy to maximize recombinant protein expression in *E. coli* is to express the protein in a host bacteria with an impaired capacity to proteolytically cleave the recombinant protein (Gottesman, p. 119-128, In *Gene Expression Technology: Methods in Enzymology* vol. 185, Academic Press, San Diego, CA, 1990. Another

10 strategy is to alter the nucleic acid sequence of the nucleic acid to be inserted into an expression vector so that the individual codons for each amino acid are those preferentially utilized in *E. coli* (Wada *et al.*, 1992, *Nucleic Acids Res.* 20:2111-2118). Such alteration of nucleic acid sequences of the invention can be carried out by standard DNA synthesis techniques.

15 In another embodiment, the expression vector is a yeast expression vector. Examples of vectors for expression in yeast *S. cerevisiae* include pYepSec1 (Baldari *et al.*, 1987, *EMBO J.* 6:229-234), pMFa (Kurjan and Herskowitz, 1982, *Cell* 30:933-943), pJRY88 (Schultz *et al.*, 1987, *Gene* 54:113-123), pYES2 (Invitrogen Corporation, San Diego, CA), and pPicZ (Invitrogen Corp, San Diego, CA).

20 Alternatively, the expression vector is a baculovirus expression vector. Baculovirus vectors available for expression of proteins in cultured insect cells (*e.g.*, Sf 9 cells) include the pAc series (Smith *et al.*, 1983, *Mol. Cell Biol.* 3:2156-2165) and the pVL series (Lucklow and Summers, 1989, *Virology* 170:31-39).

In yet another embodiment, a nucleic acid of the invention is expressed in

25 mammalian cells using a mammalian expression vector. Examples of mammalian expression vectors include pCDM8 (Seed, 1987, *Nature* 329:840) and pMT2PC (Kaufman *et al.*, 1987, *EMBO J.* 6:187-195). When used in mammalian cells, the expression vector's control functions are often provided by viral regulatory elements. For example, commonly used promoters are derived from polyoma, Adenovirus 2,

30 cytomegalovirus and Simian Virus 40. For other suitable expression systems for both prokaryotic and eukaryotic cells see chapters 16 and 17 of Sambrook *et al.*, *supra*.

In another embodiment, the recombinant mammalian expression vector is capable of directing expression of the nucleic acid preferentially in a particular cell type (*e.g.*, tissue-specific regulatory elements are used to express the nucleic acid). Tissue-specific regulatory elements are known in the art. Non-limiting examples of suitable

5 tissue-specific promoters include the albumin promoter (liver-specific; Pinkert *et al.*, 1987, *Genes Dev.* 1:268-277), lymphoid-specific promoters (Calame and Eaton, 1988, *Adv. Immunol.* 43:235-275), in particular promoters of T cell receptors (Winoto and Baltimore, 1989, *EMBO J.* 8:729-733) and immunoglobulins (Banerji *et al.*, 1983, *Cell* 33:729-740; Queen and Baltimore, 1983, *Cell* 33:741-748), neuron-specific promoters

10 (*e.g.*, the neurofilament promoter; Byrne and Ruddle, 1989, *Proc. Natl. Acad. Sci. USA* 86:5473-5477), pancreas-specific promoters (Edlund *et al.*, 1985, *Science* 230:912-916), and mammary gland-specific promoters (*e.g.*, milk whey promoter; U.S. Patent No. 4,873,316 and European Application Publication No. 264,166). Developmentally-regulated promoters are also encompassed, for example the murine hox promoters

15 (Kessel and Gruss, 1990, *Science* 249:374-379) and the α-fetoprotein promoter (Camper and Tilghman, 1989, *Genes Dev.* 3:537-546).

The invention further provides a recombinant expression vector comprising a DNA molecule of the invention cloned into the expression vector in an antisense orientation. That is, the DNA molecule is operably linked to a regulatory

20 sequence in a manner which allows for expression (by transcription of the DNA molecule) of an RNA molecule which is antisense to the mRNA encoding a polypeptide of the invention. Regulatory sequences operably linked to a nucleic acid cloned in the antisense orientation can be chosen which direct the continuous expression of the antisense RNA molecule in a variety of cell types, for instance viral promoters and/or

25 enhancers, or regulatory sequences can be chosen which direct constitutive, tissue-specific or cell type specific expression of antisense RNA. The antisense expression vector can be in the form of a recombinant plasmid, phagemid, or attenuated virus in which antisense nucleic acids are produced under the control of a high efficiency regulatory region, the activity of which can be determined by the cell type into which the

30 vector is introduced. For a discussion of the regulation of gene expression using antisense genes see Weintraub *et al.*, 1986, *Trends in Genetics*, Vol. 1(1).

Another aspect of the invention pertains to host cells into which a recombinant expression vector of the invention has been introduced. The terms "host cell" and "recombinant host cell" are used interchangeably herein. It is understood that such terms refer not only to the particular subject cell but to the progeny or potential progeny of such a cell. Because certain modifications may occur in succeeding generations due to either mutation or environmental influences, such progeny may not, in fact, be identical to the parent cell, but are still included within the scope of the term as used herein.

A host cell can be any prokaryotic (*e.g.*, *E. coli*) or eukaryotic cell (*e.g.*, insect cells, yeast or mammalian cells).

Vector DNA can be introduced into prokaryotic or eukaryotic cells via conventional transformation or transfection techniques. As used herein, the terms "transformation" and "transfection" are intended to refer to a variety of art-recognized techniques for introducing foreign nucleic acid into a host cell, including calcium phosphate or calcium chloride co-precipitation, DEAE-dextran-mediated transfection, lipofection, or electroporation. Suitable methods for transforming or transfecting host cells can be found in Sambrook, *et al.* (*supra*), and other laboratory manuals.

For stable transfection of mammalian cells, it is known that, depending upon the expression vector and transfection technique used, only a small fraction of cells may integrate the foreign DNA into their genome. In order to identify and select these integrants, a gene that encodes a selectable marker (*e.g.*, for resistance to antibiotics) is generally introduced into the host cells along with the gene of interest. Preferred selectable markers include those which confer resistance to drugs, such as G418, hygromycin and methotrexate. Cells stably transfected with the introduced nucleic acid can be identified by drug selection (*e.g.*, cells that have incorporated the selectable marker will survive, while the other cells die).

A host cell of the invention, such as a prokaryotic or eukaryotic host cell in culture, can be used to produce a marker protein or a segment thereof. Accordingly, the invention further provides methods for producing a marker protein or a segment thereof using the host cells of the invention. In one embodiment, the method comprises culturing the host cell of the invention (into which a recombinant expression vector encoding a marker protein or a segment thereof has been introduced) in a suitable medium such that the is produced. In another embodiment, the method further

comprises isolating the marker protein or a segment thereof from the medium or the host cell.

The host cells of the invention can also be used to produce nonhuman transgenic animals. For example, in one embodiment, a host cell of the invention is a fertilized oocyte or an embryonic stem cell into which a sequences encoding a marker protein or a segment thereof have been introduced. Such host cells can then be used to create non-human transgenic animals in which exogenous sequences encoding a marker protein of the invention have been introduced into their genome or homologous recombinant animals in which endogenous gene(s) encoding a marker protein have been altered. Such animals are useful for studying the function and/or activity of the marker protein and for identifying and/or evaluating modulators of marker protein. As used herein, a "transgenic animal" is a non-human animal, preferably a mammal, more preferably a rodent such as a rat or mouse, in which one or more of the cells of the animal includes a transgene. Other examples of transgenic animals include non-human primates, sheep, dogs, cows, goats, chickens, amphibians, etc. A transgene is exogenous DNA which is integrated into the genome of a cell from which a transgenic animal develops and which remains in the genome of the mature animal, thereby directing the expression of an encoded gene product in one or more cell types or tissues of the transgenic animal. As used herein, an "homologous recombinant animal" is a non-human animal, preferably a mammal, more preferably a mouse, in which an endogenous gene has been altered by homologous recombination between the endogenous gene and an exogenous DNA molecule introduced into a cell of the animal, e.g., an embryonic cell of the animal, prior to development of the animal.

A transgenic animal of the invention can be created by introducing a nucleic acid encoding a marker protein into the male pronuclei of a fertilized oocyte, e.g., by microinjection, retroviral infection, and allowing the oocyte to develop in a pseudopregnant female foster animal. Intronic sequences and polyadenylation signals can also be included in the transgene to increase the efficiency of expression of the transgene. A tissue-specific regulatory sequence(s) can be operably linked to the transgene to direct expression of the polypeptide of the invention to particular cells. Methods for generating transgenic animals via embryo manipulation and microinjection, particularly animals such as mice, have become conventional in the art and are described, for example, in U.S. Patent Nos. 4,736,866 and 4,870,009, U.S. Patent No.

4,873,191 and in Hogan, *Manipulating the Mouse Embryo*, Cold Spring Harbor

Laboratory Press, Cold Spring Harbor, N.Y., 1986. Similar methods are used for

production of other transgenic animals. A transgenic founder animal can be identified

based upon the presence of the transgene in its genome and/or expression of mRNA

5   encoding the transgene in tissues or cells of the animals. A transgenic founder animal

can then be used to breed additional animals carrying the transgene. Moreover,

transgenic animals carrying the transgene can further be bred to other transgenic animals

carrying other transgenes.

To create an homologous recombinant animal, a vector is prepared which

10   contains at least a portion of a gene encoding a marker protein into which a deletion,

addition or substitution has been introduced to thereby alter, *e.g.*, functionally disrupt,

the gene. In a preferred embodiment, the vector is designed such that, upon homologous

recombination, the endogenous gene is functionally disrupted (*i.e.*, no longer encodes a

functional protein; also referred to as a "knock out" vector). Alternatively, the vector

15   can be designed such that, upon homologous recombination, the endogenous gene is

mutated or otherwise altered but still encodes functional protein (*e.g.*, the upstream

regulatory region can be altered to thereby alter the expression of the endogenous

protein). In the homologous recombination vector, the altered portion of the gene is

flanked at its 5' and 3' ends by additional nucleic acid of the gene to allow for

20   homologous recombination to occur between the exogenous gene carried by the vector

and an endogenous gene in an embryonic stem cell. The additional flanking nucleic acid

sequences are of sufficient length for successful homologous recombination with the

endogenous gene. Typically, several kilobases of flanking DNA (both at the 5' and 3'

ends) are included in the vector (see, *e.g.*, Thomas and Capecchi, 1987, *Cell* 51:503 for a

25   description of homologous recombination vectors). The vector is introduced into an

embryonic stem cell line (*e.g.*, by electroporation) and cells in which the introduced

gene has homologously recombined with the endogenous gene are selected (see, *e.g.*, Li

et al., 1992, *Cell* 69:915). The selected cells are then injected into a blastocyst of an

animal (*e.g.*, a mouse) to form aggregation chimeras (see, *e.g.*, Bradley,

30   *Teratocarcinomas and Embryonic Stem Cells: A Practical Approach*, Robertson, Ed.,

IRL, Oxford, 1987, pp. 113-152). A chimeric embryo can then be implanted into a

suitable pseudopregnant female foster animal and the embryo brought to term. Progeny

harboring the homologously recombined DNA in their germ cells can be used to breed

animals in which all cells of the animal contain the homologously recombined DNA by germline transmission of the transgene. Methods for constructing homologous recombination vectors and homologous recombinant animals are described further in Bradley (1991) *Current Opinion in Bio/Technology* 2:823-829 and in PCT Publication

5   NOS. WO 90/11354, WO 91/01140, WO 92/0968, and WO 93/04169.

In another embodiment, transgenic non-human animals can be produced which contain selected systems which allow for regulated expression of the transgene. One example of such a system is the *cre/loxP* recombinase system of bacteriophage P1. For a description of the *cre/loxP* recombinase system, see, *e.g.*, Lakso *et al.* (1992) *Proc.*

10 *Natl. Acad. Sci. USA* 89:6232-6236. Another example of a recombinase system is the FLP recombinase system of *Saccharomyces cerevisiae* (O'Gorman *et al.*, 1991, *Science* 251:1351-1355). If a *cre/loxP* recombinase system is used to regulate expression of the transgene, animals containing transgenes encoding both the *Cre* recombinase and a selected protein are required. Such animals can be provided through the construction of

15 "double" transgenic animals, *e.g.*, by mating two transgenic animals, one containing a transgene encoding a selected protein and the other containing a transgene encoding a recombinase.

Clones of the non-human transgenic animals described herein can also be produced according to the methods described in Wilmut *et al.* (1997) *Nature* 385:810-

20 813 and PCT Publication NOS. WO 97/07668 and WO 97/07669.

IV. Pharmaceutical Compositions

The nucleic acid molecules, polypeptides, and antibodies (also referred to herein as "active compounds") of the invention can be incorporated into pharmaceutical

25 compositions suitable for administration. Such compositions typically comprise the nucleic acid molecule, protein, or antibody and a pharmaceutically acceptable carrier. As used herein the language "pharmaceutically acceptable carrier" is intended to include any and all solvents, dispersion media, coatings, antibacterial and antifungal agents, isotonic and absorption delaying agents, and the like, compatible with pharmaceutical

30 administration. The use of such media and agents for pharmaceutically active substances is well known in the art. Except insofar as any conventional media or agent is incompatible with the active compound, use thereof in the compositions is

contemplated. Supplementary active compounds can also be incorporated into the compositions.

The invention includes methods for preparing pharmaceutical compositions for modulating the expression or activity of a marker nucleic acid or

5   protein . Such methods comprise formulating a pharmaceutically acceptable carrier with an agent which modulates expression or activity of a marker nucleic acid or protein. Such compositions can further include additional active agents. Thus, the invention further includes methods for preparing a pharmaceutical composition by formulating a pharmaceutically acceptable carrier with an agent which modulates expression or

10   activity of a marker nucleic acid or protein and one or more additional active compounds.

The invention also provides methods (also referred to herein as "screening assays") for identifying modulators, *i.e.,* candidate or test compounds or agents (*e.g.,* peptides, peptidomimetics, peptoids, small molecules or other drugs) which

15   (a) bind to the marker, or (b) have a modulatory (*e.g.,* stimulatory or inhibitory) effect on the activity of the marker or, more specifically, (c) have a modulatory effect on the interactions of the marker with one or more of its natural substrates (*e.g.,* peptide, protein, hormone, co-factor, or nucleic acid), or (d) have a modulatory effect on the expression of the marker. Such assays typically comprise a reaction between the marker

20   and one or more assay components. The other components may be either the test compound itself, or a combination of test compound and a natural binding partner of the marker.

The test compounds of the present invention may be obtained from any available source, including systematic libraries of natural and/or synthetic compounds.

25   Test compounds may also be obtained by any of the numerous approaches in combinatorial library methods known in the art, including: biological libraries; peptoid libraries (libraries of molecules having the functionalities of peptides, but with a novel, non-peptide backbone which are resistant to enzymatic degradation but which nevertheless remain bioactive; see, *e.g.,* Zuckermann *et al.,* 1994, *J. Med. Chem.*

30   37:2678-85); spatially addressable parallel solid phase or solution phase libraries; synthetic library methods requiring deconvolution; the 'one-bead one-compound' library method; and synthetic library methods using affinity chromatography selection. The biological library and peptoid library approaches are limited to peptide libraries, while

the other four approaches are applicable to peptide, non-peptide oligomer or small molecule libraries of compounds (Lam, 1997, *Anticancer Drug Des.* 12:145).

Examples of methods for the synthesis of molecular libraries can be found in the art, for example in: DeWitt *et al.* (1993) *Proc. Natl. Acad. Sci. U.S.A.*
90:6909; Erb *et al.* (1994) *Proc. Natl. Acad. Sci. USA* 91:11422; Zuckermann *et al.* (1994). *J. Med. Chem.* 37:2678; Cho *et al.* (1993) *Science* 261:1303; Carrell *et al.* (1994) *Angew. Chem. Int. Ed. Engl.* 33:2059; Carell *et al.* (1994) *Angew. Chem. Int. Ed. Engl.* 33:2061; and in Gallop *et al.* (1994) *J. Med. Chem.* 37:1233.

Libraries of compounds may be presented in solution (*e.g.*, Houghten, 1992, *Biotechniques* 13:412-421), or on beads (Lam, 1991, *Nature* 354:82-84), chips (Fodor, 1993, *Nature* 364:555-556), bacteria and/or spores, (Ladner, USP 5,223,409), plasmids (Cull *et al*, 1992, *Proc Natl Acad Sci USA* 89:1865-1869) or on phage (Scott -and Smith, 1990, *Science* 249:386-390; Devlin, 1990, *Science* 249:404-406; Cwirla *et al*, 1990, *Proc. Natl. Acad. Sci.* 87:6378-6382; Felici, 1991, *J. Mol. Biol.* 222:301-310; Ladner, *supra.*).

In one embodiment, the invention provides assays for screening candidate or test compounds which are substrates of a protein encoded by or corresponding to a marker or biologically active portion thereof. In another embodiment, the invention provides assays for screening candidate or test compounds which bind to a protein encoded by or corresponding to a marker or biologically active portion thereof. Determining the ability of the test compound to directly bind to a protein can be accomplished, for example, by coupling the compound with a radioisotope or enzymatic label such that binding of the compound to the marker can be determined by detecting the labeled marker compound in a complex. For example, compounds (*e.g.*, marker substrates) can be labeled with ^{125}I, ^{35}S, ^{14}C, or ^{3}H, either directly or indirectly, and the radioisotope detected by direct counting of radioemission or by scintillation counting. Alternatively, assay components can be enzymatically labeled with, for example, horseradish peroxidase, alkaline phosphatase, or luciferase, and the enzymatic label detected by determination of conversion of an appropriate substrate to product.

In another embodiment, the invention provides assays for screening candidate or test compounds which modulate the expression of a marker or the activity of a protein encoded by or corresponding to a marker, or a biologically active portion

thereof. In all likelihood, the protein encoded by or corresponding to the marker can, *in vivo*, interact with one or more molecules, such as but not limited to, peptides, proteins, hormones, cofactors and nucleic acids. For the purposes of this discussion, such cellular and extracellular molecules are referred to herein as "binding partners" or marker

5   "substrate".

One necessary embodiment of the invention in order to facilitate such screening is the use of a protein encoded by or corresponding to marker to identify the protein's natural *in vivo* binding partners. There are many ways to accomplish this which are known to one skilled in the art. One example is the use of the marker protein

10   as "bait protein" in a two-hybrid assay or three-hybrid assay (see, *e.g.*, U.S. Patent No. 5,283,317; Zervos *et al*, 1993, *Cell* 72:223-232; Madura *et al*, 1993, *J. Biol. Chem.* 268:12046-12054; Bartel *et al*, 1993, *Biotechniques* 14:920-924; Iwabuchi *et al*, 1993 *Oncogene* 8:1693-1696; Brent WO94/10300) in order to identify other proteins which bind to or interact with the marker (binding partners) and, therefore, are possibly

15   involved in the natural function of the marker. Such marker binding partners are also likely to be involved in the propagation of signals by the marker protein or downstream elements of a marker protein-mediated signaling pathway. Alternatively, such marker protein binding partners may also be found to be inhibitors of the marker protein.

The two-hybrid system is based on the modular nature of most

20   transcription factors, which consist of separable DNA-binding and activation domains. Briefly, the assay utilizes two different DNA constructs. In one construct, the gene that encodes a marker protein fused to a gene encoding the DNA binding domain of a known transcription factor (*e.g.*, GAL-4). In the other construct, a DNA sequence, from a library of DNA sequences, that encodes an unidentified protein ("prey" or "sample") is

25   fused to a gene that codes for the activation domain of the known transcription factor. If the "bait" and the "prey" proteins are able to interact, *in vivo*, forming a marker-dependent complex, the DNA-binding and activation domains of the transcription factor are brought into close proximity. This proximity allows transcription of a reporter gene (*e.g.*, LacZ) which is operably linked to a transcriptional regulatory site responsive to

30   the transcription factor. Expression of the reporter gene can be readily detected and cell colonies containing the functional transcription factor can be isolated and used to obtain the cloned gene which encodes the protein which interacts with the marker protein.

In a further embodiment, assays may be devised through the use of the invention for the purpose of identifying compounds which modulate (*e.g.,* affect either positively or negatively) interactions between a marker protein and its substrates and/or binding partners. Such compounds can include, but are not limited to, molecules such as antibodies, peptides, hormones, oligonucleotides, nucleic acids, and analogs thereof. Such compounds may also be obtained from any available source, including systematic libraries of natural and/or synthetic compounds. The preferred assay components for use in this embodiment is an cervical cancer marker protein identified herein, the known binding partner and/or substrate of same, and the test compound. Test compounds can be supplied from any source.

The basic principle of the assay systems used to identify compounds that interfere with the interaction between the marker protein and its binding partner involves preparing a reaction mixture containing the marker protein and its binding partner under conditions and for a time sufficient to allow the two products to interact and bind, thus forming a complex. In order to test an agent for inhibitory activity, the reaction mixture is prepared in the presence and absence of the test compound. The test compound can be initially included in the reaction mixture, or can be added at a time subsequent to the addition of the marker protein and its binding partner. Control reaction mixtures are incubated without the test compound or with a placebo. The formation of any complexes between the marker protein and its binding partner is then detected. The formation of a complex in the control reaction, but less or no such formation in the reaction mixture containing the test compound, indicates that the compound interferes with the interaction of the marker protein and its binding partner. Conversely, the formation of more complex in the presence of compound than in the control reaction indicates that the compound may enhance interaction of the marker protein and its binding partner.

The assay for compounds that interfere with the interaction of the marker protein with its binding partner may be conducted in a heterogeneous or homogeneous format. Heterogeneous assays involve anchoring either the marker protein or its binding partner onto a solid phase and detecting complexes anchored to the solid phase at the end of the reaction. In homogeneous assays, the entire reaction is carried out in a liquid phase. In either approach, the order of addition of reactants can be varied to obtain different information about the compounds being tested. For example, test compounds

that interfere with the interaction between the marker proteins and the binding partners (*e.g.*, by competition) can be identified by conducting the reaction in the presence of the test substance, *i.e.*, by adding the test substance to the reaction mixture prior to or simultaneously with the marker and its interactive binding partner. Alternatively, test

5   compounds that disrupt preformed complexes, *e.g.*, compounds with higher binding constants that displace one of the components from the complex, can be tested by adding the test compound to the reaction mixture after complexes have been formed. The various formats are briefly described below.

In a heterogeneous assay system, either the marker protein or its binding

10   partner is anchored onto a solid surface or matrix, while the other corresponding non-anchored component may be labeled, either directly or indirectly. In practice, microtitre plates are often utilized for this approach. The anchored species can be immobilized by a number of methods, either non-covalent or covalent, that are typically well known to one who practices the art. Non-covalent attachment can often be accomplished simply by

15   coating the solid surface with a solution of the marker protein or its binding partner and drying. Alternatively, an immobilized antibody specific for the assay component to be anchored can be used for this purpose. Such surfaces can often be prepared in advance and stored.

In related embodiments, a fusion protein can be provided which adds a

20   domain that allows one or both of the assay components to be anchored to a matrix. For example, glutathione-S-transferase/marker fusion proteins or glutathione-S-transferase/binding partner can be adsorbed onto glutathione sepharose beads (Sigma Chemical, St. Louis, MO) or glutathione derivatized microtiter plates, which are then combined with the test compound or the test compound and either the non-adsorbed

25   marker or its binding partner, and the mixture incubated under conditions conducive to complex formation (*e.g.*, physiological conditions). Following incubation, the beads or microtiter plate wells are washed to remove any unbound assay components, the immobilized complex assessed either directly or indirectly, for example, as described above. Alternatively, the complexes can be dissociated from the matrix, and the level of

30   marker binding or activity determined using standard techniques.

Other techniques for immobilizing proteins on matrices can also be used in the screening assays of the invention. For example, either a marker protein or a marker protein binding partner can be immobilized utilizing conjugation of biotin and

streptavidin. Biotinylated marker protein or target molecules can be prepared from biotin-NHS (N-hydroxy-succinimide) using techniques known in the art (*e.g.*, biotinylation kit, Pierce Chemicals, Rockford, IL), and immobilized in the wells of streptavidin-coated 96 well plates (Pierce Chemical). In certain embodiments, the
5   protein-immobilized surfaces can be prepared in advance and stored.

In order to conduct the assay, the corresponding partner of the immobilized assay component is exposed to the coated surface with or without the test compound. After the reaction is complete, unreacted assay components are removed (*e.g.*, by washing) and any complexes formed will remain immobilized on the solid
10   surface. The detection of complexes anchored on the solid surface can be accomplished in a number of ways. Where the non-immobilized component is pre-labeled, the detection of label immobilized on the surface indicates that complexes were formed. Where the non-immobilized component is not pre-labeled, an indirect label can be used to detect complexes anchored on the surface; *e.g.*, using a labeled antibody specific for
15   the initially non-immobilized species (the antibody, in turn, can be directly labeled or indirectly labeled with, *e.g.*, a labeled anti-Ig antibody). Depending upon the order of addition of reaction components, test compounds which modulate (inhibit or enhance) complex formation or which disrupt preformed complexes can be detected.

In an alternate embodiment of the invention, a homogeneous assay may
20   be used. This is typically a reaction, analogous to those mentioned above, which is conducted in a liquid phase in the presence or absence of the test compound. The formed complexes are then separated from unreacted components, and the amount of complex formed is determined. As mentioned for heterogeneous assay systems, the order of addition of reactants to the liquid phase can yield information about which test
25   compounds modulate (inhibit or enhance) complex formation and which disrupt preformed complexes.

In such a homogeneous assay, the reaction products may be separated from unreacted assay components by any of a number of standard techniques, including but not limited to: differential centrifugation, chromatography, electrophoresis and
30   immunoprecipitation. In differential centrifugation, complexes of molecules may be separated from uncomplexed molecules through a series of centrifugal steps, due to the different sedimentation equilibria of complexes based on their different sizes and densities (see, for example, Rivas, G., and Minton, A.P., *Trends Biochem Sci* 1993

Aug;18(8):284-7). Standard chromatographic techniques may also be utilized to separate complexed molecules from uncomplexed ones. For example, gel filtration chromatography separates molecules based on size, and through the utilization of an appropriate gel filtration resin in a column format, for example, the relatively larger

5   complex may be separated from the relatively smaller uncomplexed components. Similarly, the relatively different charge properties of the complex as compared to the uncomplexed molecules may be exploited to differentially separate the complex from the remaining individual reactants, for example through the use of ion-exchange chromatography resins. Such resins and chromatographic techniques are well known to

10   one skilled in the art (see, *e.g.*, Heegaard, 1998, *J Mol. Recognit.* 11:141-148; Hage and Tweed, 1997, *J. Chromatogr. B. Biomed. Sci. Appl.*, 699:499-525). Gel electrophoresis may also be employed to separate complexed molecules from unbound species (see, *e.g.*, Ausubel *et al* (eds.), In: Current Protocols in Molecular Biology, J. Wiley & Sons, New York. 1999). In this technique, protein or nucleic acid complexes are separated based on

15   size or charge, for example. In order to maintain the binding interaction during the electrophoretic process, nondenaturing gels in the absence of reducing agent are typically preferred, but conditions appropriate to the particular interactants will be well known to one skilled in the art. Immunoprecipitation is another common technique utilized for the isolation of a protein-protein complex from solution (see, *e.g.*, Ausubel *et*

20   *al* (eds.), In: Current Protocols in Molecular Biology, J. Wiley & Sons, New York. 1999). In this technique, all proteins binding to an antibody specific to one of the binding molecules are precipitated from solution by conjugating the antibody to a polymer bead that may be readily collected by centrifugation. The bound assay components are released from the beads (through a specific proteolysis event or other

25   technique well known in the art which will not disturb the protein-protein interaction in the complex), and a second immunoprecipitation step is performed, this time utilizing antibodies specific for the correspondingly different interacting assay component. In this manner, only formed complexes should remain attached to the beads. Variations in complex formation in both the presence and the absence of a test compound can be

30   compared, thus offering information about the ability of the compound to modulate interactions between the marker protein and its binding partner.

Also within the scope of the present invention are methods for direct detection of interactions between the marker protein and its natural binding partner and/or a test compound in a homogeneous or heterogeneous assay system without further sample manipulation. For example, the technique of fluorescence energy transfer

5   may be utilized (see, e.g., Lakowicz et al, U.S. Patent No. 5,631,169; Stavrianopoulos et al, U.S. Patent No. 4,868,103). Generally, this technique involves the addition of a fluorophore label on a first 'donor' molecule (e.g., marker or test compound) such that its emitted fluorescent energy will be absorbed by a fluorescent label on a second, 'acceptor' molecule (e.g., marker or test compound), which in turn is able to fluoresce

10   due to the absorbed energy. Alternately, the 'donor' protein molecule may simply utilize the natural fluorescent energy of tryptophan residues. Labels are chosen that emit different wavelengths of light, such that the 'acceptor' molecule label may be differentiated from that of the 'donor'. Since the efficiency of energy transfer between the labels is related to the distance separating the molecules, spatial relationships

15   between the molecules can be assessed. In a situation in which binding occurs between the molecules, the fluorescent emission of the 'acceptor' molecule label in the assay should be maximal. An FET binding event can be conveniently measured through standard fluorometric detection means well known in the art (e.g., using a fluorimeter). A test substance which either enhances or hinders participation of one of the species in

20   the preformed complex will result in the generation of a signal variant to that of background. In this way, test substances that modulate interactions between a marker and its binding partner can be identified in controlled assays.

In another embodiment, modulators of marker expression are identified in a method wherein a cell is contacted with a candidate compound and the expression

25   of marker mRNA or protein in the cell, is determined. The level of expression of marker mRNA or protein in the presence of the candidate compound is compared to the level of expression of marker mRNA or protein in the absence of the candidate compound. The candidate compound can then be identified as a modulator of marker expression based on this comparison. For example, when expression of marker mRNA

30   or protein is greater (statistically significantly greater) in the presence of the candidate compound than in its absence, the candidate compound is identified as a stimulator of marker mRNA or protein expression. Conversely, when expression of marker mRNA or protein is less (statistically significantly less) in the presence of the candidate compound

than in its absence, the candidate compound is identified as an inhibitor of marker mRNA or protein expression. The level of marker mRNA or protein expression in the cells can be determined by methods described herein for detecting marker mRNA or protein.

In another aspect, the invention pertains to a combination of two or more of the assays described herein. For example, a modulating agent can be identified using a cell-based or a cell free assay, and the ability of the agent to modulate the activity of a marker protein can be further confirmed *in vivo*, *e.g.*, in a whole animal model for cellular transformation and/or tumorigenesis.

This invention further pertains to novel agents identified by the above-described screening assays. Accordingly, it is within the scope of this invention to further use an agent identified as described herein in an appropriate animal model. For example, an agent identified as described herein (*e.g.*, a marker modulating agent, an antisense marker nucleic acid molecule, a marker-specific antibody, or a marker-binding partner) can be used in an animal model to determine the efficacy, toxicity, or side effects of treatment with such an agent. Alternatively, an agent identified as described herein can be used in an animal model to determine the mechanism of action of such an agent. Furthermore, this invention pertains to uses of novel agents identified by the above-described screening assays for treatments as described herein.

It is understood that appropriate doses of small molecule agents and protein or polypeptide agents depends upon a number of factors within the knowledge of the ordinarily skilled physician, veterinarian, or researcher. The dose(s) of these agents will vary, for example, depending upon the identity, size, and condition of the subject or sample being treated, further depending upon the route by which the composition is to be administered, if applicable, and the effect which the practitioner desires the agent to have upon the nucleic acid or polypeptide of the invention. Exemplary doses of a small molecule include milligram or microgram amounts per kilogram of subject or sample weight (*e.g.* about 1 microgram per kilogram to about 500 milligrams per kilogram, about 100 micrograms per kilogram to about 5 milligrams per kilogram, or about 1 microgram per kilogram to about 50 micrograms per kilogram). Exemplary doses of a protein or polypeptide include gram, milligram or microgram amounts per kilogram of subject or sample weight (*e.g.* about 1 microgram per kilogram to about 5 grams per kilogram, about 100 micrograms per kilogram to about 500 milligrams per kilogram, or

about 1 milligram per kilogram to about 50 milligrams per kilogram). It is furthermore understood that appropriate doses of one of these agents depend upon the potency of the agent with respect to the expression or activity to be modulated. Such appropriate doses can be determined using the assays described herein. When one or more of these agents

5   is to be administered to an animal (e.g. a human) in order to modulate expression or activity of a polypeptide or nucleic acid of the invention, a physician, veterinarian, or researcher can, for example, prescribe a relatively low dose at first, subsequently increasing the dose until an appropriate response is obtained. In addition, it is understood that the specific dose level for any particular animal subject will depend

10   upon a variety of factors including the activity of the specific agent employed, the age, body weight, general health, gender, and diet of the subject, the time of administration, the route of administration, the rate of excretion, any drug combination, and the degree of expression or activity to be modulated.

A pharmaceutical composition of the invention is formulated to be

15   compatible with its intended route of administration. Examples of routes of administration include parenteral, e.g., intravenous, intradermal, subcutaneous, oral (e.g., inhalation), transdermal (topical), transmucosal, and rectal administration. Solutions or suspensions used for parenteral, intradermal, or subcutaneous application can include the following components: a sterile diluent such as water for injection, saline

20   solution, fixed oils, polyethylene glycols, glycerine, propylene glycol or other synthetic solvents; antibacterial agents such as benzyl alcohol or methyl parabens; antioxidants such as ascorbic acid or sodium bisulfite; chelating agents such as ethylenediamine-tetraacetic acid; buffers such as acetates, citrates or phosphates and agents for the adjustment of tonicity such as sodium chloride or dextrose. pH can be adjusted with

25   acids or bases, such as hydrochloric acid or sodium hydroxide. The parenteral preparation can be enclosed in ampules, disposable syringes or multiple dose vials made of glass or plastic.

Pharmaceutical compositions suitable for injectable use include sterile aqueous solutions (where water soluble) or dispersions and sterile powders for the

30   extemporaneous preparation of sterile injectable solutions or dispersions. For intravenous administration, suitable carriers include physiological saline, bacteriostatic water, Cremophor EL (BASF; Parsippany, NJ) or phosphate buffered saline (PBS). In all cases, the composition must be sterile and should be fluid to the extent that easy

syringability exists. It must be stable under the conditions of manufacture and storage and must be preserved against the contaminating action of microorganisms such as bacteria and fungi. The carrier can be a solvent or dispersion medium containing, for example, water, ethanol, polyol (for example, glycerol, propylene glycol, and liquid

5   polyethylene glycol, and the like), and suitable mixtures thereof. The proper fluidity can be maintained, for example, by the use of a coating such as lecithin, by the maintenance of the required particle size in the case of dispersion and by the use of surfactants. Prevention of the action of microorganisms can be achieved by various antibacterial and antifungal agents, for example, parabens, chlorobutanol, phenol, ascorbic acid,

10   thimerosal, and the like. In many cases, it will be preferable to include isotonic agents, for example, sugars, polyalcohols such as mannitol, sorbitol, or sodium chloride in the composition. Prolonged absorption of the injectable compositions can be brought about by including in the composition an agent which delays absorption, for example, aluminum monostearate and gelatin.

15   Sterile injectable solutions can be prepared by incorporating the active compound (*e.g.*, a polypeptide or antibody) in the required amount in an appropriate solvent with one or a combination of ingredients enumerated above, as required, followed by filtered sterilization. Generally, dispersions are prepared by incorporating the active compound into a sterile vehicle which contains a basic dispersion medium,

20   and then incorporating the required other ingredients from those enumerated above. In the case of sterile powders for the preparation of sterile injectable solutions, the preferred methods of preparation are vacuum drying and freeze-drying which yields a powder of the active ingredient plus any additional desired ingredient from a previously sterile-filtered solution thereof.

25   Oral compositions generally include an inert diluent or an edible carrier. They can be enclosed in gelatin capsules or compressed into tablets. For the purpose of oral therapeutic administration, the active compound can be incorporated with excipients and used in the form of tablets, troches, or capsules. Oral compositions can also be prepared using a fluid carrier for use as a mouthwash, wherein the compound in the fluid

30   carrier is applied orally and swished and expectorated or swallowed.

               Pharmaceutically compatible binding agents, and/or adjuvant materials can be included as part of the composition. The tablets, pills, capsules, troches, and the like can contain any of the following ingredients, or compounds of a similar nature: a

binder such as microcrystalline cellulose, gum tragacanth or gelatin; an excipient such as starch or lactose, a disintegrating agent such as alginic acid, Primogel, or corn starch; a lubricant such as magnesium stearate or Sterotes; a glidant such as colloidal silicon dioxide; a sweetening agent such as sucrose or saccharin; or a flavoring agent such as peppermint, methyl salicylate, or orange flavoring.

For administration by inhalation, the compounds are delivered in the form of an aerosol spray from a pressurized container or dispenser which contains a suitable propellant, *e.g.*, a gas such as carbon dioxide, or a nebulizer.

Systemic administration can also be by transmucosal or transdermal means. For transmucosal or transdermal administration, penetrants appropriate to the barrier to be permeated are used in the formulation. Such penetrants are generally known in the art, and include, for example, for transmucosal administration, detergents, bile salts, and fusidic acid derivatives. Transmucosal administration can be accomplished through the use of nasal sprays or suppositories. For transdermal administration, the active compounds are formulated into ointments, salves, gels, or creams as generally known in the art.

The compounds can also be prepared in the form of suppositories (*e.g.*, with conventional suppository bases such as cocoa butter and other glycerides) or retention enemas for rectal delivery.

In one embodiment, the active compounds are prepared with carriers that will protect the compound against rapid elimination from the body, such as a controlled release formulation, including implants and microencapsulated delivery systems. Biodegradable, biocompatible polymers can be used, such as ethylene vinyl acetate, polyanhydrides, polyglycolic acid, collagen, polyorthoesters, and polylactic acid. Methods for preparation of such formulations will be apparent to those skilled in the art. The materials can also be obtained commercially from Alza Corporation and Nova Pharmaceuticals, Inc. Liposomal suspensions (including liposomes having monoclonal antibodies incorporated therein or thereon) can also be used as pharmaceutically acceptable carriers. These can be prepared according to methods known to those skilled in the art, for example, as described in U.S. Patent No. 4,522,811.

It is especially advantageous to formulate oral or parenteral compositions in dosage unit form for ease of administration and uniformity of dosage. Dosage unit form as used herein refers to physically discrete units suited as unitary dosages for the

subject to be treated; each unit containing a predetermined quantity of active compound calculated to produce the desired therapeutic effect in association with the required pharmaceutical carrier. The specification for the dosage unit forms of the invention are dictated by and directly dependent on the unique characteristics of the active compound

5 and the particular therapeutic effect to be achieved, and the limitations inherent in the art of compounding such an active compound for the treatment of individuals.

For antibodies, the preferred dosage is 0.1 mg/kg to 100 mg/kg of body weight (generally 10 mg/kg to 20 mg/kg). If the antibody is to act in the brain, a dosage of 50 mg/kg to 100 mg/kg is usually appropriate. Generally, partially human antibodies

10 and fully human antibodies have a longer half-life within the human body than other antibodies. Accordingly, lower dosages and less frequent administration is often possible. Modifications such as lipidation can be used to stabilize antibodies and to enhance uptake and tissue penetration (*e.g.*, into the cervical epithelium). A method for lipidation of antibodies is described by Cruikshank *et al.* (1997) *J. Acquired Immune*

15 *Deficiency Syndromes and Human Retrovirology* 14:193.

The invention also provides vaccine compositions for the prevention and/or treatment of cervical cancer. The invention provides cervical cancer vaccine compositions in which a protein of a marker of Table 1, or a combination of proteins of the markers of Table 1, are introduced into a subject in order to stimulate an immune

20 response against the cervical cancer. The invention also provides cervical cancer vaccine compositions in which a gene expression construct, which expresses a marker or fragment of a marker identified in Table 1, is introduced into the subject such that a protein or fragment of a protein encoded by a marker of Table 1 is produced by transfected cells in the subject at a higher than normal level and elicits an immune

25 response.

In one embodiment, a cervical cancer vaccine is provided and employed as an immunotherapeutic agent for the prevention of cervical cancer. In another embodiment, a cervical cancer vaccine is provided and employed as an immunotherapeutic agent for the treatment of cervical cancer.

30 By way of example, a cervical cancer vaccine comprised of the proteins of the markers of Table 1, may be employed for the prevention and/or treatment of cervical cancer in a subject by administering the vaccine by a variety of routes, *e.g.*, intradermally, subcutaneously, or intramuscularly. In addition, the cervical cancer

vaccine can be administered together with adjuvants and/or immunomodulators to boost the activity of the vaccine and the subject's response. In one embodiment, devices and/or compositions containing the vaccine, suitable for sustained or intermittent release could be, implanted in the body or topically applied thereto for the relatively slow

5   release of such materials into the body. The cervical cancer vaccine can be introduced along with immunomodulatory compounds, which can alter the type of immune response produced in order to produce a response which will be more effective in eliminating the cancer.

In another embodiment, a cervical cancer vaccine comprised of an

10   expression construct of the markers of Table 1, may be introduced by injection into muscle or by coating onto microprojectiles and using a device designed for the purpose to fire the projectiles at high speed into the skin. The cells of the subject will then express the protein(s) or fragments of proteins of the markers of Table 1 and induce an immune

15   response. In addition, the cervical cancer vaccine may be introduced along with expression constructs for immunomodulatory molecules, such as cytokines, which may increase the immune response or modulate the type of immune response produced in order to produce a response which will be more effective in eliminating the cancer.

The marker nucleic acid molecules can be inserted into vectors and used

20   as gene therapy vectors. Gene therapy vectors can be delivered to a subject by, for example, intravenous injection, local administration (U.S. Patent 5,328,470), or by stereotactic injection (see, e.g., Chen et al., 1994, *Proc. Natl. Acad. Sci. USA* 91:3054-3057). The pharmaceutical preparation of the gene therapy vector can include the gene therapy vector in an acceptable diluent, or can comprise a slow release matrix in which

25   the gene delivery vehicle is imbedded. Alternatively, where the complete gene delivery vector can be produced intact from recombinant cells, *e.g.* retroviral vectors, the pharmaceutical preparation can include one or more cells which produce the gene delivery system.

The pharmaceutical compositions can be included in a container, pack, or

30   dispenser together with instructions for administration.

V. Predictive Medicine

The present invention pertains to the field of predictive medicine in which diagnostic assays, prognostic assays, pharmacogenomics, and monitoring clinical trails are used for prognostic (predictive) purposes to thereby treat an individual prophylactically. Accordingly, one aspect of the present invention relates to diagnostic assays for determining the level of expression of one or more marker proteins or nucleic acids, in order to determine whether an individual is at risk of developing cervical cancer. Such assays can be used for prognostic or predictive purposes to thereby prophylactically treat an individual prior to the onset of the cancer.

Yet another aspect of the invention pertains to monitoring the influence of agents (*e.g.*, drugs or other compounds administered either to inhibit cervical cancer or to treat or prevent any other disorder {*i.e.* in order to understand any cervical carcinogenic effects that such treatment may have}) on the expression or activity of a marker of the invention in clinical trials. These and other agents are described in further detail in the following sections.

A. Diagnostic Assays

An exemplary method for detecting the presence or absence of a marker protein or nucleic acid in a biological sample involves obtaining a biological sample (*e.g.* a cervical-associated body fluid) from a test subject and contacting the biological sample with a compound or an agent capable of detecting the polypeptide or nucleic acid (*e.g.*, mRNA, genomic DNA, or cDNA). The detection methods of the invention can thus be used to detect mRNA, protein, cDNA, or genomic DNA, for example, in a biological sample *in vitro* as well as *in vivo*. For example, *in vitro* techniques for detection of mRNA include Northern hybridizations and *in situ* hybridizations. *In vitro* techniques for detection of a marker protein include enzyme linked immunosorbent assays (ELISAs), Western blots, immunoprecipitations and immunofluorescence. *In vitro* techniques for detection of genomic DNA include Southern hybridizations. Furthermore, *in vivo* techniques for detection of a marker protein include introducing into a subject a labeled antibody directed against the protein or fragment thereof. For example, the antibody can be labeled with a radioactive marker whose presence and location in a subject can be detected by standard imaging techniques.

A general principle of such diagnostic and prognostic assays involves preparing a sample or reaction mixture that may contain a marker, and a probe, under appropriate conditions and for a time sufficient to allow the marker and probe to interact and bind, thus forming a complex that can be removed and/or detected in the reaction
5   mixture. These assays can be conducted in a variety of ways.

For example, one method to conduct such an assay would involve anchoring the marker or probe onto a solid phase support, also referred to as a substrate, and detecting target marker/probe complexes anchored on the solid phase at the end of the reaction. In one embodiment of such a method, a sample from a subject, which is to
10   be assayed for presence and/or concentration of marker, can be anchored onto a carrier or solid phase support. In another embodiment, the reverse situation is possible, in which the probe can be anchored to a solid phase and a sample from a subject can be allowed to react as an unanchored component of the assay.

There are many established methods for anchoring assay components to
15   a solid phase. These include, without limitation, marker or probe molecules which are immobilized through conjugation of biotin and streptavidin. Such biotinylated assay components can be prepared from biotin-NHS (N-hydroxy-succinimide) using techniques known in the art (e.g., biotinylation kit, Pierce Chemicals, Rockford, IL), and immobilized in the wells of streptavidin-coated 96 well plates (Pierce Chemical). In
20   certain embodiments, the surfaces with immobilized assay components can be prepared in advance and stored.

Other suitable carriers or solid phase supports for such assays include any material capable of binding the class of molecule to which the marker or probe belongs. Well-known supports or carriers include, but are not limited to, glass, polystyrene,
25   nylon, polypropylene, nylon, polyethylene, dextran, amylases, natural and modified celluloses, polyacrylamides, gabbros, and magnetite.

In order to conduct assays with the above mentioned approaches, the non-immobilized component is added to the solid phase upon which the second component is anchored. After the reaction is complete, uncomplexed components may
30   be removed (e.g., by washing) under conditions such that any complexes formed will remain immobilized upon the solid phase. The detection of marker/probe complexes anchored to the solid phase can be accomplished in a number of methods outlined herein.

In a preferred embodiment, the probe, when it is the unanchored assay component, can be labeled for the purpose of detection and readout of the assay, either directly or indirectly, with detectable labels discussed herein and which are well-known to one skilled in the art.

It is also possible to directly detect marker/probe complex formation without further manipulation or labeling of either component (marker or probe), for example by utilizing the technique of fluorescence energy transfer (see, for example, Lakowicz et al., U.S. Patent No. 5,631,169; Stavrianopoulos, et al., U.S. Patent No. 4,868,103). A fluorophore label on the first, 'donor' molecule is selected such that, upon excitation with incident light of appropriate wavelength, its emitted fluorescent energy will be absorbed by a fluorescent label on a second 'acceptor' molecule, which in turn is able to fluoresce due to the absorbed energy. Alternately, the 'donor' protein molecule may simply utilize the natural fluorescent energy of tryptophan residues. Labels are chosen that emit different wavelengths of light, such that the 'acceptor' molecule label may be differentiated from that of the 'donor'. Since the efficiency of energy transfer between the labels is related to the distance separating the molecules, spatial relationships between the molecules can be assessed. In a situation in which binding occurs between the molecules, the fluorescent emission of the 'acceptor' molecule label in the assay should be maximal. An FET binding event can be conveniently measured through standard fluorometric detection means well known in the art (e.g., using a fluorimeter).

In another embodiment, determination of the ability of a probe to recognize a marker can be accomplished without labeling either assay component (probe or marker) by utilizing a technology such as real-time Biomolecular Interaction Analysis (BIA) (see, e.g., Sjolander, S. and Urbaniczky, C., 1991, Anal. Chem. 63:2338-2345 and Szabo et al., 1995, Curr. Opin. Struct. Biol. 5:699-705). As used herein, "BIA" or "surface plasmon resonance" is a technology for studying biospecific interactions in real time, without labeling any of the interactants (e.g., BIAcore). Changes in the mass at the binding surface (indicative of a binding event) result in alterations of the refractive index of light near the surface (the optical phenomenon of surface plasmon resonance (SPR)), resulting in a detectable signal which can be used as an indication of real-time reactions between biological molecules.

Alternatively, in another embodiment, analogous diagnostic and prognostic assays can be conducted with marker and probe as solutes in a liquid phase. In such an assay, the complexed marker and probe are separated from uncomplexed components by any of a number of standard techniques, including but not limited to:

5   differential centrifugation, chromatography, electrophoresis and immunoprecipitation. In differential centrifugation, marker/probe complexes may be separated from uncomplexed assay components through a series of centrifugal steps, due to the different sedimentation equilibria of complexes based on their different sizes and densities (see, for example, Rivas, G., and Minton, A.P., 1993, *Trends Biochem Sci.* 18(8):284-7).

10   Standard chromatographic techniques may also be utilized to separate complexed molecules from uncomplexed ones. For example, gel filtration chromatography separates molecules based on size, and through the utilization of an appropriate gel filtration resin in a column format, for example, the relatively larger complex may be separated from the relatively smaller uncomplexed components. Similarly, the relatively

15   different charge properties of the marker/probe complex as compared to the uncomplexed components may be exploited to differentiate the complex from uncomplexed components, for example through the utilization of ion-exchange chromatography resins. Such resins and chromatographic techniques are well known to one skilled in the art (see, *e.g.*, Heegaard, N.H., 1998, *J. Mol. Recognit.* Winter 11(1-

20   6):141-8; Hage, D.S., and Tweed, S.A. *J Chromatogr B Biomed Sci Appl* 1997 Oct 10;699(1-2):499-525). Gel electrophoresis may also be employed to separate complexed assay components from unbound components (see, *e.g.*, Ausubel *et al.*, ed., *Current Protocols in Molecular Biology*, John Wiley & Sons, New York, 1987-1999). In this technique, protein or nucleic acid complexes are separated based on size or charge, for

25   example. In order to maintain the binding interaction during the electrophoretic process, non-denaturing gel matrix materials and conditions in the absence of reducing agent are typically preferred. Appropriate conditions to the particular assay and components thereof will be well known to one skilled in the art.

In a particular embodiment, the level of marker mRNA can be

30   determined both by *in situ* and by *in vitro* formats in a biological sample using methods known in the art. The term "biological sample" is intended to include tissues, cells, biological fluids and isolates thereof, isolated from a subject, as well as tissues, cells and fluids present within a subject. Many expression detection methods use isolated RNA.

For *in vitro* methods, any RNA isolation technique that does not select against the isolation of mRNA can be utilized for the purification of RNA from cervical cells (see, *e.g.*, Ausubel *et al.*, ed., *Current Protocols in Molecular Biology*, John Wiley & Sons, New York 1987-1999). Additionally, large numbers of tissue samples can readily be

5   processed using techniques well known to those of skill in the art, such as, for example, the single-step RNA isolation process of Chomczynski (1989, U.S. Patent No. 4,843,155).

The isolated mRNA can be used in hybridization or amplification assays that include, but are not limited to, Southern or Northern analyses, polymerase chain

10   reaction analyses and probe arrays. One preferred diagnostic method for the detection of mRNA levels involves contacting the isolated mRNA with a nucleic acid molecule (probe) that can hybridize to the mRNA encoded by the gene being detected. The nucleic acid probe can be, for example, a full-length cDNA, or a portion thereof, such as an oligonucleotide of at least 7, 15, 30, 50, 100, 250 or 500 nucleotides in length and

15   sufficient to specifically hybridize under stringent conditions to a mRNA or genomic DNA encoding a marker of the present invention. Other suitable probes for use in the diagnostic assays of the invention are described herein. Hybridization of an mRNA with the probe indicates that the marker in question is being expressed.

In one format, the mRNA is immobilized on a solid surface and contacted

20   with a probe, for example by running the isolated mRNA on an agarose gel and transferring the mRNA from the gel to a membrane, such as nitrocellulose. In an alternative format, the probe(s) are immobilized on a solid surface and the mRNA is contacted with the probe(s), for example, in an Affymetrix gene chip array. A skilled artisan can readily adapt known mRNA detection methods for use in detecting the level

25   of mRNA encoded by the markers of the present invention.

An alternative method for determining the level of mRNA marker in a sample involves the process of nucleic acid amplification, *e.g.*, by rtPCR (the experimental embodiment set forth in Mullis, 1987, U.S. Patent No. 4,683,202), ligase chain reaction (Barany, 1991, *Proc. Natl. Acad. Sci. USA*, 88:189-193), self sustained

30   sequence replication (Guatelli *et al.*, 1990, *Proc. Natl. Acad. Sci. USA* 87:1874-1878), transcriptional amplification system (Kwoh *et al.*, 1989, *Proc. Natl. Acad. Sci. USA* 86:1173-1177), Q-Beta Replicase (Lizardi *et al.*, 1988, *Bio/Technology* 6:1197), rolling circle replication (Lizardi *et al.*, U.S. Patent No. 5,854,033) or any other nucleic acid

amplification method, followed by the detection of the amplified molecules using techniques well known to those of skill in the art. These detection schemes are especially useful for the detection of nucleic acid molecules if such molecules are present in very low numbers. As used herein, amplification primers are defined as being

5   a pair of nucleic acid molecules that can anneal to 5' or 3' regions of a gene (plus and minus strands, respectively, or vice-versa) and contain a short region in between. In general, amplification primers are from about 10 to 30 nucleotides in length and flank a region from about 50 to 200 nucleotides in length. Under appropriate conditions and with appropriate reagents, such primers permit the amplification of a nucleic acid

10   molecule comprising the nucleotide sequence flanked by the primers.

For *in situ* methods, mRNA does not need to be isolated from the cervical cells prior to detection. In such methods, a cell or tissue sample is prepared/processed using known histological methods. The sample is then immobilized on a support, typically a glass slide, and then contacted with a probe that can hybridize to mRNA that

15   encodes the marker.

As an alternative to making determinations based on the absolute expression level of the marker, determinations may be based on the normalized expression level of the marker. Expression levels are normalized by correcting the absolute expression level of a marker by comparing its expression to the expression of a

20   gene that is not a marker, *e.g.*, a housekeeping gene that is constitutively expressed. Suitable genes for normalization include housekeeping genes such as the actin gene, or epithelial cell-specific genes. This normalization allows the comparison of the expression level in one sample, *e.g.*, a patient sample, to another sample, *e.g.*, a non-cervical cancer sample, or between samples from different sources.

25   Alternatively, the expression level can be provided as a relative expression level. To determine a relative expression level of a marker, the level of expression of the marker is determined for 10 or more samples of normal versus cancer cell isolates, preferably 50 or more samples, prior to the determination of the expression level for the sample in question. The mean expression level of each of the genes assayed

30   in the larger number of samples is determined and this is used as a baseline expression level for the marker. The expression level of the marker determined for the test sample (absolute level of expression) is then divided by the mean expression value obtained for that marker. This provides a relative expression level.

Preferably, the samples used in the baseline determination will be from cervical cancer or from non-cervical cancer cells of cervical tissue. The choice of the cell source is dependent on the use of the relative expression level. Using expression found in normal tissues as a mean expression score aids in validating whether the marker assayed is cervical specific (versus normal cells). In addition, as more data is accumulated, the mean expression value can be revised, providing improved relative expression values based on accumulated data. Expression data from cervical cells provides a means for grading the severity of the cervical cancer state.

In another embodiment of the present invention, a marker protein is detected. A preferred agent for detecting marker protein of the invention is an antibody capable of binding to such a protein or a fragment thereof, preferably an antibody with a detectable label. Antibodies can be polyclonal, or more preferably, monoclonal. An intact antibody, or a fragment or derivative thereof (*e.g.*, Fab or F(ab')$_2$) can be used. The term "labeled", with regard to the probe or antibody, is intended to encompass direct labeling of the probe or antibody by coupling (*i.e.*, physically linking) a detectable substance to the probe or antibody, as well as indirect labeling of the probe or antibody by reactivity with another reagent that is directly labeled. Examples of indirect labeling include detection of a primary antibody using a fluorescently labeled secondary antibody and end-labeling of a DNA probe with biotin such that it can be detected with fluorescently labeled streptavidin.

Proteins from cervical cells can be isolated using techniques that are well known to those of skill in the art. The protein isolation methods employed can, for example, be such as those described in Harlow and Lane (Harlow and Lane, 1988, *Antibodies: A Laboratory Manual*, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, New York).

A variety of formats can be employed to determine whether a sample contains a protein that binds to a given antibody. Examples of such formats include, but are not limited to, enzyme immunoassay (EIA), radioimmunoassay (RIA), Western blot analysis and enzyme linked immunoabsorbant assay (ELISA). A skilled artisan can readily adapt known protein/antibody detection methods for use in determining whether cervical cells express a marker of the present invention.

In one format, antibodies, or antibody fragments or derivatives, can be used in methods such as Western blots or immunofluorescence techniques to detect the expressed proteins. In such uses, it is generally preferable to immobilize either the antibody or proteins on a solid support. Suitable solid phase supports or carriers include any support capable of binding an antigen or an antibody. Well-known supports or carriers include glass, polystyrene, polypropylene, polyethylene, dextran, nylon, amylases, natural and modified celluloses, polyacrylamides, gabbros, and magnetite.

One skilled in the art will know many other suitable carriers for binding antibody or antigen, and will be able to adapt such support for use with the present invention. For example, protein isolated from cervical cells can be run on a polyacrylamide gel electrophoresis and immobilized onto a solid phase support such as nitrocellulose. The support can then be washed with suitable buffers followed by treatment with the detectably labeled antibody. The solid phase support can then be washed with the buffer a second time to remove unbound antibody. The amount of bound label on the solid support can then be detected by conventional means.

The invention also encompasses kits for detecting the presence of a marker protein or nucleic acid in a biological sample (*e.g.*, cervical smear). Such kits can be used to determine if a subject is suffering from or is at increased risk of developing cervical cancer. For example, the kit can comprise a labeled compound or agent capable of detecting a marker protein or nucleic acid in a biological sample and means for determining the amount of the protein or mRNA in the sample (*e.g.*, an antibody which binds the protein or a fragment thereof, or an oligonucleotide probe which binds to DNA or mRNA encoding the protein). Kits can also include instructions for interpreting the results obtained using the kit.

For antibody-based kits, the kit can comprise, for example: (1) a first antibody (*e.g.*, attached to a solid support) which binds to a marker protein; and, optionally, (2) a second, different antibody which binds to either the protein or the first antibody and is conjugated to a detectable label.

For oligonucleotide-based kits, the kit can comprise, for example: (1) an oligonucleotide, *e.g.*, a detectably labeled oligonucleotide, which hybridizes to a nucleic acid sequence encoding a marker protein or (2) a pair of primers useful for amplifying a marker nucleic acid molecule. The kit can also comprise, *e.g.*, a buffering agent, a preservative, or a protein stabilizing agent. The kit can further comprise components

necessary for detecting the detectable label (*e.g.*, an enzyme or a substrate). The kit can also contain a control sample or a series of control samples which can be assayed and compared to the test sample. Each component of the kit can be enclosed within an individual container and all of the various containers can be within a single package, along with instructions for interpreting the results of the assays performed using the kit.

B. Pharmacogenomics

The markers of the invention are also useful as pharmacogenomic markers. As used herein, a "pharmacogenomic marker" is an objective biochemical marker whose expression level correlates with a specific clinical drug response or susceptibility in a patient (see, *e.g.*, McLeod *et al.* (1999) *Eur. J. Cancer* 35(12): 1650-1652). The presence or quantity of the pharmacogenomic marker expression is related to the predicted response of the patient and more particularly the patient's tumor to therapy with a specific drug or class of drugs. By assessing the presence or quantity of the expression of one or more pharmacogenomic markers in a patient, a drug therapy which is most appropriate for the patient, or which is predicted to have a greater degree of success, may be selected. For example, based on the presence or quantity of RNA or protein encoded by specific tumor markers in a patient, a drug or course of treatment may be selected that is optimized for the treatment of the specific tumor likely to be present in the patient. The use of pharmacogenomic markers therefore permits selecting or designing the most appropriate treatment for each cancer patient without trying different drugs or regimes.

Another aspect of pharmacogenomics deals with genetic conditions that alters the way the body acts on drugs. These pharmacogenetic conditions can occur either as rare defects or as polymorphisms. For example, glucose-6-phosphate dehydrogenase (G6PD) deficiency is a common inherited enzymopathy in which the main clinical complication is hemolysis after ingestion of oxidant drugs (anti-malarials, sulfonamides, analgesics, nitrofurans) and consumption of fava beans.

As an illustrative embodiment, the activity of drug metabolizing enzymes is a major determinant of both the intensity and duration of drug action. The discovery of genetic polymorphisms of drug metabolizing enzymes (*e.g.*, N-acetyltransferase 2 (NAT 2) and cytochrome P450 enzymes CYP2D6 and CYP2C19) has provided an explanation as to why some patients do not obtain the expected drug effects or show

exaggerated drug response and serious toxicity after taking the standard and safe dose of a drug. These polymorphisms are expressed in two phenotypes in the population, the extensive metabolizer (EM) and poor metabolizer (PM). The prevalence of PM is different among different populations. For example, the gene coding for CYP2D6 is

5   highly polymorphic and several mutations have been identified in PM, which all lead to the absence of functional CYP2D6. Poor metabolizers of CYP2D6 and CYP2C19 quite frequently experience exaggerated drug response and side effects when they receive standard doses. If a metabolite is the active therapeutic moiety, a PM will show no therapeutic response, as demonstrated for the analgesic effect of codeine mediated by its

10   CYP2D6-formed metabolite morphine. The other extreme are the so called ultra-rapid metabolizers who do not respond to standard doses. Recently, the molecular basis of ultra-rapid metabolism has been identified to be due to CYP2D6 gene amplification.

Thus, the level of expression of a marker of the invention in an individual can be determined to thereby select appropriate agent(s) for therapeutic or prophylactic

15   treatment of the individual. In addition, pharmacogenetic studies can be used to apply genotyping of polymorphic alleles encoding drug-metabolizing enzymes to the identification of an individual's drug responsiveness phenotype. This knowledge, when applied to dosing or drug selection, can avoid adverse reactions or therapeutic failure and thus enhance therapeutic or prophylactic efficiency when treating a subject with a

20   modulator of expression of a marker of the invention.

C. Monitoring Clinical Trials

Monitoring the influence of agents (*e.g.*, drug compounds) on the level of expression of a marker of the invention can be applied not only in basic drug screening,

25   but also in clinical trials. For example, the effectiveness of an agent to affect marker expression can be monitored in clinical trials of subjects receiving treatment for cervical cancer. In a preferred embodiment, the present invention provides a method for monitoring the effectiveness of treatment of a subject with an agent (*e.g.*, an agonist, antagonist, peptidomimetic, protein, peptide, nucleic acid, small molecule, or other drug

30   candidate) comprising the steps of (i) obtaining a pre-administration sample from a subject prior to administration of the agent; (ii) detecting the level of expression of one or more selected markers of the invention in the pre-administration sample; (iii) obtaining one or more post-administration samples from the subject; (iv) detecting the

level of expression of the marker(s) in the post-administration samples; (v) comparing the level of expression of the marker(s) in the pre-administration sample with the level of expression of the marker(s) in the post-administration sample or samples; and (vi) altering the administration of the agent to the subject accordingly. For example,

5   increased expression of the marker gene(s) during the course of treatment may indicate ineffective dosage and the desirability of increasing the dosage. Conversely, decreased expression of the marker gene(s) may indicate efficacious treatment and no need to change dosage.

10   D. Electronic Apparatus Readable Media and Arrays

Electronic apparatus readable media comprising a marker of the present invention is also provided. As used herein, "electronic apparatus readable media" refers to any suitable medium for storing, holding or containing data or information that can be read and accessed directly by an electronic apparatus. Such media can include, but are

15   not limited to: magnetic storage media, such as floppy discs, hard disc storage medium, and magnetic tape; optical storage media such as compact disc; electronic storage media such as RAM, ROM, EPROM, EEPROM and the like; general hard disks and hybrids of these categories such as magnetic/optical storage media. The medium is adapted or configured for having recorded thereon a marker of the present invention.

20   As used herein, the term "electronic apparatus" is intended to include any suitable computing or processing apparatus or other device configured or adapted for storing data or information. Examples of electronic apparatus suitable for use with the present invention include stand-alone computing apparatus; networks, including a local area network (LAN), a wide area network (WAN) Internet, Intranet, and Extranet;

25   electronic appliances such as a personal digital assistants (PDAs), cellular phone, pager and the like; and local and distributed processing systems.

As used herein, "recorded" refers to a process for storing or encoding information on the electronic apparatus readable medium. Those skilled in the art can readily adopt any of the presently known methods for recording information on known

30   media to generate manufactures comprising the markers of the present invention.

A variety of software programs and formats can be used to store the marker information of the present invention on the electronic apparatus readable medium. For example, the marker nucleic acid sequence can be represented in a word

processing text file, formatted in commercially-available software such as WordPerfect and MicroSoft Word, or represented in the form of an ASCII file, stored in a database application, such as DB2, Sybase, Oracle, or the like, as well as in other forms. Any number of data processor structuring formats (e.g., text file or database) may be

5   employed in order to obtain or create a medium having recorded thereon the markers of the present invention.

By providing the markers of the invention in readable form, one can routinely access the marker sequence information for a variety of purposes. For example, one skilled in the art can use the nucleotide or amino acid sequences of the

10   present invention in readable form to compare a target sequence or target structural motif with the sequence information stored within the data storage means. Search means are used to identify fragments or regions of the sequences of the invention which match a particular target sequence or target motif.

The present invention therefore provides a medium for holding

15   instructions for performing a method for determining whether a subject has cervical cancer or a pre-disposition to cervical cancer, wherein the method comprises the steps of determining the presence or absence of a marker and based on the presence or absence of the marker, determining whether the subject has cervical cancer or a pre-disposition to cervical cancer and/or recommending a particular treatment for cervical cancer or pre-

20   cervical cancer condition.

The present invention further provides in an electronic system and/or in a network, a method for determining whether a subject has cervical cancer or a pre-disposition to cervical cancer associated with a marker wherein the method comprises the steps of determining the presence or absence of the marker, and based on the

25   presence or absence of the marker, determining whether the subject has cervical cancer or a pre-disposition to cervical cancer, and/or recommending a particular treatment for the cervical cancer or pre-cervical cancer condition. The method may further comprise the step of receiving phenotypic information associated with the subject and/or acquiring from a network phenotypic information associated with the subject.

30   The present invention also provides in a network, a method for determining whether a subject has cervical cancer or a pre-disposition to cervical cancer associated with a marker, said method comprising the steps of receiving information associated with the marker receiving phenotypic information associated with the subject,

acquiring information from the network corresponding to the marker and/or cervical cancer, and based on one or more of the phenotypic information, the marker, and the acquired information, determining whether the subject has a cervical cancer or a pre-disposition to cervical cancer. The method may further comprise the step of

5   recommending a particular treatment for the cervical cancer or pre-cervical cancer condition.

The present invention also provides a business method for determining whether a subject has cervical cancer or a pre-disposition to cervical cancer, said method comprising the steps of receiving information associated with the marker, receiving

10   phenotypic information associated with the subject, acquiring information from the network corresponding to the marker and/or cervical cancer, and based on one or more of the phenotypic information, the marker, and the acquired information, determining whether the subject has cervical cancer or a pre-disposition to cervical cancer. The method may further comprise the step of recommending a particular treatment for the

15   cervical cancer or pre-cervical cancer condition.

The invention also includes an array comprising a marker of the present invention. The array can be used to assay expression of one or more genes in the array. In one embodiment, the array can be used to assay gene expression in a tissue to ascertain tissue specificity of genes in the array. In this manner, up to about 7600 genes

20   can be simultaneously assayed for expression. This allows a profile to be developed showing a battery of genes specifically expressed in one or more tissues.

In addition to such qualitative determination, the invention allows the quantitation of gene expression. Thus, not only tissue specificity, but also the level of expression of a battery of genes in the tissue is ascertainable. Thus, genes can be

25   grouped on the basis of their tissue expression *per se* and level of expression in that tissue. This is useful, for example, in ascertaining the relationship of gene expression between or among tissues. Thus, one tissue can be perturbed and the effect on gene expression in a second tissue can be determined. In this context, the effect of one cell type on another cell type in response to a biological stimulus can be determined. Such a

30   determination is useful, for example, to know the effect of cell-cell interaction at the level of gene expression. If an agent is administered therapeutically to treat one cell type but has an undesirable effect on another cell type, the invention provides an assay to determine the molecular basis of the undesirable effect and thus provides the

opportunity to co-administer a counteracting agent or otherwise treat the undesired effect. Similarly, even within a single cell type, undesirable biological effects can be determined at the molecular level. Thus, the effects of an agent on expression of other than the target gene can be ascertained and counteracted.

5   In another embodiment, the array can be used to monitor the time course of expression of one or more genes in the array. This can occur in various biological contexts, as disclosed herein, for example development of cervical cancer, progression of cervical cancer, and processes, such a cellular transformation associated with cervical cancer.

10   The array is also useful for ascertaining the effect of the expression of a gene on the expression of other genes in the same cell or in different cells. This provides, for example, for a selection of alternate molecular targets for therapeutic intervention if the ultimate or downstream target cannot be regulated.

The array is also useful for ascertaining differential expression patterns of

15   one or more genes in normal and abnormal cells. This provides a battery of genes that could serve as a molecular target for diagnosis or therapeutic intervention.

E. Surrogate Markers

The markers of the invention may serve as surrogate markers for one or

20   more disorders or disease states or for conditions leading up to disease states, and in particular, cervical cancer. As used herein, a "surrogate marker" is an objective biochemical marker which correlates with the absence or presence of a disease or disorder, or with the progression of a disease or disorder (e.g., with the presence or absence of a tumor). The presence or quantity of such markers is independent of the

25   disease. Therefore, these markers may serve to indicate whether a particular course of treatment is effective in lessening a disease state or disorder. Surrogate markers are of particular use when the presence or extent of a disease state or disorder is difficult to assess through standard methodologies (e.g., early stage tumors), or when an assessment of disease progression is desired before a potentially dangerous clinical endpoint is

30   reached (e.g., an assessment of cardiovascular disease may be made using cholesterol levels as a surrogate marker, and an analysis of HIV infection may be made using HIV RNA levels as a surrogate marker, well in advance of the undesirable clinical outcomes of myocardial infarction or fully-developed AIDS). Examples of the use of surrogate

markers in the art include: Koomen *et al.* (2000) *J. Mass. Spectrom.* 35: 258-264; and

James (1994) *AIDS Treatment News Archive* 209.

The markers of the invention are also useful as pharmacodynamic

markers. As used herein, a "pharmacodynamic marker" is an objective biochemical

5   marker which correlates specifically with drug effects. The presence or quantity of a

pharmacodynamic marker is not related to the disease state or disorder for which the

drug is being administered; therefore, the presence or quantity of the marker is indicative

of the presence or activity of the drug in a subject. For example, a pharmacodynamic

marker may be indicative of the concentration of the drug in a biological tissue, in that

10   the marker is either expressed or transcribed or not expressed or transcribed in that tissue

in relationship to the level of the drug. In this fashion, the distribution or uptake of the

drug may be monitored by the pharmacodynamic marker. Similarly, the presence or

quantity of the pharmacodynamic marker may be related to the presence or quantity of

the metabolic product of a drug, such that the presence or quantity of the marker is

15   indicative of the relative breakdown rate of the drug *in vivo*. Pharmacodynamic markers

are of particular use in increasing the sensitivity of detection of drug effects, particularly

when the drug is administered in low doses. Since even a small amount of a drug may

be sufficient to activate multiple rounds of marker transcription or expression, the

amplified marker may be in a quantity which is more readily detectable than the drug

20   itself. Also, the marker may be more easily detected due to the nature of the marker

itself; for example, using the methods described herein, antibodies may be employed in

an immune-based detection system for a protein marker, or marker-specific radiolabeled

probes may be used to detect a mRNA marker. Furthermore, the use of a

pharmacodynamic marker may offer mechanism-based prediction of risk due to drug

25   treatment beyond the range of possible direct observations. Examples of the use of

pharmacodynamic markers in the art include: Matsuda *et al.* US 6,033,862; Hattis *et al.*

(1991) *Env. Health Perspect.* 90: 229-238; Schentag (1999) *Am. J. Health-Syst. Pharm.*

56 Suppl. 3: S21-S24; and Nicolau (1999) *Am, J. Health-Syst. Pharm.* 56 Suppl. 3: S16-

S20.

30

VI.   Experimental Protocol

A. Identification of clones

Cervical tumor specific cDNA clones were identified by transcription profiling using mRNA from 12 cervical tumors, 5 CIN III, 5 CIN I and 12 normal cervical tissues. The subtracted libraries were constructed using mRNA from at least three independent normal ectocervix, B-lymphocytes, T-lymphocytes and other white blood cells (in activated and resting states) as drivers and four independent stage 1B cervical tumors or four independent C1N III cervical samples as testers. The top up-regulated clones in tumors or C1N III cervical tissues, as determined by proprietary statistical analysis methods, were selected. The clusters in which the selected clones belong were blasted against both public and proprietary sequence databases in order to identify other EST sequences or clusters with significant overlap. Thus, contiguous EST sequences and/or clusters were assembled into full-length genes.

An identification of protein sequence corresponding to the clone was accomplished by obtaining one of the following:

a) a direct match between the protein sequence and at least one EST sequence in one of its 6 possible translations;

b) a direct match between the nucleotide sequence for the mRNA corresponding to the protein sequence and at least one EST sequence;

c) a match between the protein sequence and a contiguous assembly (contig) of the EST sequences with other available EST sequences in the databases in one of its 6 possible translations; or

d) a match between the nucleotide sequence for the mRNA corresponding to the protein sequence and a contiguous assembly of the EST sequences with other available EST sequences in the databases in one of its 6 possible translations.

VII. Summary of the Data

Tables 1-3 list the markers obtained using the foregoing protocol. The tables provide the name of the gene corresponding to the marker ("Gene Name"), the sequence listing identifier of the cDNA sequence of a nucleotide transcript encoded by or corresponding to the marker ("SEQ ID NO (nts)"), the sequence listing identifier of the amino acid sequence of a protein encoded by the nucleotide transcript ("SEQ ID NO

(AAs)"), and the location of the protein coding sequence within the cDNA sequence ("CDS").

Table 1 lists all of the markers of the invention which are over-expressed in cervical cancer cells compared to normal (*i.e.,* non-cancerous) cervical cells. Table 2 lists newly-identified nucleotide and amino acid sequences useful as cervical cancer markers. Table 3 lists newly-identified nucleotide sequences useful as cervical cancer markers.

Other Embodiments

Those skilled in the art will recognize, or be able to ascertain using no more than routine experimentation, many equivalents to the specific embodiments of the invention described herein. Such equivalents are intended to be encompassed by the following claims:

What is claimed:

1.   An isolated nucleic acid molecule comprising a nucleotide sequence selected from the group consisting of SEQ ID NOs: 1, 3, 5, 7, 143, 145, 147, 149, 151, 167, 203, 217, 231, 233, 51, 65, 67, 68, 100, and 153.

2.   A vector which contains the nucleic acid molecule of claim 1.

3.   A host cell which contains the nucleic acid molecule of claim 1.

4.   A method of assessing whether a patient is afflicted with cervical cancer, the method comprising comparing:

      a) the level of expression of a marker in a patient sample, wherein the marker is selected from Table 1; and

      b) the normal level of expression of the marker in a control non-cervical cancer sample,

      wherein a significant increase in the level of expression of the marker in the patient sample and the normal level is an indication that the patient is afflicted with cervical cancer.

5.   An isolated polypeptide which is encoded by a nucleic acid molecule comprising a nucleotide sequence selected from the group consisting of SEQ ID NOs: 1, 3, 5, 7, 143, 145, 147, 149, 151, 167, 203, 217, 231, and 233.

6.   An antibody which selectively binds to the polypeptide of claim 5.

7.   An isolated polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NOs: 2, 4, 6, 8, 144, 146, 148, 150, 152, 168, 204, 218, 232, and 234.

8.   An antibody which selectively binds to the polypeptide of claim 7.

SEQUENCE LISTING

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Ser Lys Lys Gln Lys Lys Lys Arg Lys Thr Ser Ser Ser Lys His Asp
           35                  40                  45
Val Ser Ala His His Asp Leu Asn Ile Asp Gln Ser Gln Cys Asn Glu
       50                  55                  60
Met Tyr Ile Asn Ser Ser Gln Arg Val Glu Ser Thr Val Ile Pro Glu
65                  70                  75                  80
Ser Thr Ile Met Arg Thr Leu His Ser Gly Glu Ile Thr Ser His Glu
               85                  90                  95
Gln Gly Phe Ser Val Glu Leu Glu Ser Glu Ile Ser Thr Thr Ala Asp
           100                 105                 110
Asp Cys Ser Ser Glu Val Asn Gly Cys Ser Phe Val Met Arg Thr Gly
           115                 120                 125
Lys Pro Thr Asn Leu Leu Arg Glu Glu Glu Phe Gly Val Asp Asp Ser
       130                 135                 140
Tyr Ser Glu Gln Gly Ala Gln Asp Ser Pro Thr His Leu Glu Met Met
145                 150                 155                 160
Glu Ser Glu Leu Ala Gly Lys Gln His Glu Ile Glu Glu Leu Asn Arg
               165                 170                 175
Glu Leu Glu Glu Met Arg Val Thr Tyr Gly Thr Glu Gly Leu Gln Gln
           180                 185                 190
Leu Gln Glu Phe Glu Ala Ala Ile Lys Gln Arg Asp Gly Ile Ile Thr
           195                 200                 205
Gln Leu Thr Ala Asn Leu Gln Gln Ala Arg Arg Glu Lys Asp Glu Thr
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Met Arg Glu Phe Leu Glu Leu Thr Glu Gln Ser Gln Lys Leu Gln Ile
225                 230                 235                 240
Gln Phe Gln Gln Leu Gln Ala Ser Glu Thr Leu Arg Asn Ser Thr His
               245                 250                 255
Ser Ser Thr Ala Ala Asp Leu Leu Gln Ala Lys Gln Gln Ile Leu Thr
           260                 265                 270
His Gln Gln Gln Leu Glu Glu Gln Asp His Leu Leu Glu Asp Tyr Gln
           275                 280                 285
Lys Lys Lys Glu Asp Phe Thr Met Gln Ile Ser Phe Leu Gln Glu Lys
       290                 295                 300
Ile Lys Val Tyr Glu Met Glu Gln Asp Lys Lys Val Glu Asn Ser Asn
305                 310                 315                 320
Lys Glu Glu Ile Gln Glu Lys Glu Thr Ile Ile Glu Glu Leu Asn Thr
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Lys Ile Ile Glu Glu Glu Lys Lys Thr Leu Glu Leu Lys Asp Lys Leu
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Thr Thr Ala Asp Lys Leu Leu Gly Glu Leu Gln Glu Gln Ile Val Gln
           355                 360                 365
Lys Asn Gln Glu Ile Lys Asn Met Lys Leu Glu Leu Thr Asn Ser Lys
       370                 375                 380
Gln Lys Glu Arg Gln Ser Ser Glu Glu Ile Lys Gln Leu Met Gly Thr
385                 390                 395                 400
Val Glu Glu Leu Gln Lys Arg Asn His Lys Asp Ser Gln Phe Glu Thr
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            420                     425                     430
Leu Arg Ala Glu Leu Asp Glu Met Tyr Gly Gln Gln Ile Val Gln Met
            435                     440                     445
Lys Gln Glu Leu Ile Arg Gln His Met Ala Gln Met Glu Glu Met Lys
        450                     455                     460
Thr Arg His Lys Gly Glu Met Glu Asn Ala Leu Arg Ser Tyr Ser Asn
465                     470                     475                     480
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Glu Leu Asn Ile Lys Leu Gln Asp Thr Asn Ser Gln Lys Glu Lys Leu
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Lys Glu Glu Leu Gly Leu Ile Leu Glu Glu Lys Cys Ala Leu Gln Arg
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Gln Leu Glu Asp Leu Val Glu Glu Leu Ser Phe Ser Arg Glu Gln Ile
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Gln Arg Ala Arg Gln Thr Ile Ala Glu Gln Glu Ser Lys Leu Asn Glu
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Ala His Lys Ser Leu Ser Thr Val Glu Asp Leu Lys Ala Glu Ile Val
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Ser Ala Ser Glu Ser Arg Lys Glu Leu Glu Leu Lys His Glu Ala Glu
            580                     585                     590
Val Thr Asn Tyr Lys Ile Lys Leu Glu Met Leu Glu Lys Glu Lys Asn
        595                     600                     605
Ala Val Leu Asp Arg Met Ala Glu Ser Gln Glu Ala Glu Leu Glu Arg
    610                     615                     620
Leu Arg Thr Gln Leu Leu Phe Ser His Glu Glu Glu Leu Ser Lys Leu
625                     630                     635                     640
Lys Glu Asp Leu Glu Ile Glu His Arg Ile Asn Ile Glu Lys Leu Lys
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Asp Asn Leu Gly Ile His Tyr Lys Gln Gln Ile Asp Gly Leu Gln Asn
            660                     665                     670
Glu Met Ser Gln Lys Ile Glu Thr Met Gln Phe Glu Lys Asp Asn Leu
        675                     680                     685
Ile Thr Lys Gln Asn Gln Leu Ile Leu Glu Ile Ser Lys Leu Lys Asp
    690                     695                     700
Leu Gln Gln Ser Leu Val Asn Ser Lys Ser Glu Glu Met Thr Leu Gln
705                     710                     715                     720
Ile Asn Glu Leu Gln Lys Glu Ile Glu Ile Leu Arg Gln Glu Glu Lys
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Glu Lys Gly Thr Leu Glu Gln Glu Val Gln Glu Leu Gln Leu Lys Thr
            740                     745                     750
Glu Leu Leu Glu Lys Gln Met Lys Glu Lys Glu Asn Asp Leu Gln Glu
        755                     760                     765
Lys Phe Ala Gln Leu Glu Ala Glu Asn Ser Ile Leu Lys Asp Glu Lys
    770                     775                     780
Lys Thr Leu Glu Asp Met Leu Lys Ile His Thr Pro Val Ser Gln Glu
785                     790                     795                     800
Glu Arg Leu Ile Phe Leu Asp Ser Ile Lys Ser Lys Ser Lys Asp Ser
            805                     810                     815
Val Trp Glu Lys Glu Ile Glu Ile Leu Ile Glu Glu Asn Glu Asp Leu
            820                     825                     830
Lys Gln Gln Cys Ile Gln Leu Asn Glu Glu Ile Glu Lys Gln Arg Asn
        835                     840                     845
Thr Phe Ser Phe Ala Glu Lys Asn Phe Glu Val Asn Tyr Gln Glu Leu
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Gln Glu Glu Tyr Ala Cys Leu Leu Lys Val Lys Asp Asp Leu Glu Asp
865                     870                     875                     880
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Ser Lys Asn Lys Gln Glu Leu Glu Tyr Lys Ser Lys Leu Lys Ala Leu
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Asn Glu Glu Leu His Leu Gln Arg Ile Asn Pro Thr Thr Val Lys Met
         900                 905                 910
Lys Ser Ser Val Phe Asp Glu Asp Lys Thr Phe Val Ala Glu Thr Leu
         915                 920                 925
Glu Met Gly Glu Val Val Glu Lys Asp Thr Thr Glu Leu Met Glu Lys
     930                 935                 940
Leu Glu Val Thr Lys Arg Glu Lys Leu Glu Leu Ser Gln Arg Leu Ser
945                 950                 955                 960
Asp Leu Ser Glu Gln Leu Lys Gln Lys His Gly Glu Ile Ser Phe Leu
             965                 970                 975
Asn Glu Glu Val Lys Ser Leu Lys Gln Glu Lys Glu Gln Val Ser Leu
             980                 985                 990
Arg Cys Arg Glu Leu Glu Ile Ile Ile Asn His Asn Arg Ala Glu Asn
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Val Gln Ser Cys Asp Thr Gln Val Ser Ser Leu Leu Asp Gly Val Val
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Thr Met Thr Ser Arg Gly Ala Glu Gly Ser Val Ser Lys Val Asn Lys
1025                1030                1035                1040
Ser Phe Gly Glu Glu Ser Lys Ile Met Val Glu Asp Lys Val Ser Phe
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Glu Asn Met Thr Val Gly Glu Glu Ser Lys Gln Glu Gln Leu Ile Leu
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Asp His Leu Pro Ser Val Thr Lys Glu Ser Ser Leu Arg Ala Thr Gln
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Pro Ser Glu Asn Asp Lys Leu Gln Lys Glu Leu Asn Val Leu Lys Ser
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Glu Gln Asn Asp Leu Arg Leu Gln Met Glu Ala Gln Arg Ile Cys Leu
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Ser Leu Val Tyr Ser Thr His Val Asp Gln Val Arg Glu Tyr Met Glu
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Asn Glu Lys Asp Lys Ala Leu Cys Ser Leu Lys Glu Glu Leu Ile Phe
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Ala Gln Glu Glu Lys Ile Lys Glu Leu Gln Lys Ile His Gln Leu Glu
         1155                1160                1165
Leu Gln Thr Met Lys Thr Gln Glu Thr Gly Asp Glu Gly Lys Pro Leu
     1170                1175                1180
His Leu Leu Ile Gly Lys Leu Gln Lys Ala Val Ser Glu Glu Cys Ser
1185                1190                1195                1200
Tyr Phe Leu Gln Thr Leu Cys Ser Val Leu Gly Glu Tyr Tyr Thr Pro
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Ala Leu Lys Cys Glu Val Asn Ala Glu Asp Lys Glu Asn Ser Gly Asp
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Tyr Ile Ser Glu Asn Glu Asp Pro Glu Leu Gln Asp Tyr Arg Tyr Glu
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Val Gln Asp Phe Gln Glu Asn Met His Thr Leu Leu Asn Lys Val Thr
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Glu Glu Tyr Asn Lys Leu Leu Val Leu Gln Thr Arg Leu Ser Lys Ile
1265                1270                1275                1280
Trp Gly Gln Gln Thr Asp Gly Met Lys Leu Glu Phe Gly Glu Glu Asn
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Leu Pro Lys Glu Glu Thr Glu Phe Leu Ser Ile His Ser Gln Met Thr
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Gln Asp Leu Glu Lys Thr Lys Leu Glu Glu Gln Val Gln Glu Leu Glu
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Ser Leu Ile Ser Ser Leu Gln Gln Gln Leu Lys Glu Thr Glu Gln Asn
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Glu Ser Thr Val Pro Pro Ser Leu Pro Val Asp Ser Val Val Ile Thr
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Glu Ser Asp Ala Gln Arg Thr Met Tyr Pro Gly Ser Cys Val Lys Lys
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Asn Ile Asp Gly Thr Ile Glu Phe Ser Gly Glu Phe Gly Val Lys Glu
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Glu Thr Asn Ile Val Lys Leu Leu Glu Lys Gln Tyr Gln Glu Gln Leu
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Glu Glu Glu Val Ala Lys Val Ile Val Ser Met Ser Ile Ala Phe Ala
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Gln Gln Thr Glu Leu Ser Arg Ile Ser Gly Gly Lys Glu Asn Thr Ala
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Ser Ser Lys Gln Ala His Ala Val Cys Gln Gln Glu Gln His Tyr Phe
        1475              1480              1485
Asn Glu Met Lys Leu Ser Gln Asp Gln Ile Gly Phe Gln Thr Phe Glu
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Thr Val Asp Val Lys Phe Lys Glu Glu Phe Lys Pro Leu Ser Lys Glu
1505              1510              1515              1520
Leu Gly Glu His Gly Lys Glu Ile Leu Leu Ser Asn Ser Asp Pro His
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Asp Ile Pro Glu Ser Lys Asp Cys Val Leu Thr Ile Ser Glu Glu Met
            1540              1545              1550
Phe Ser Lys Asp Lys Thr Phe Ile Val Arg Gln Ser Ile His Asp Glu
        1555              1560              1565
Ile Ser Val Ser Ser Met Asp Ala Ser Arg Gln Leu Met Leu Asn Glu
    1570              1575              1580
Glu Gln Leu Glu Asp Met Arg Gln Glu Leu Val Arg Gln Tyr Gln Glu
1585              1590              1595              1600
His Gln Gln Ala Thr Glu Leu Leu Arg Gln Ala His Met Arg Gln Met
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Leu Asn Arg Gln Leu Ala Gln Arg Ser Ser Ile Asp Asn Glu Asn Leu
        1635              1640              1645
Val Ser Glu Arg Glu Arg Val Leu Leu Glu Glu Leu Glu Ala Leu Lys
    1650              1655              1660
Gln Leu Ser Leu Ala Gly Arg Glu Lys Leu Cys Cys Glu Leu Arg Asn
1665              1670              1675              1680
Ser Ser Thr Gln Thr Gln Asn Gly Asn Glu Asn Gln Gly Glu Val Glu
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Glu Gln Thr Phe Lys Glu Lys Glu Leu Asp Arg Lys Pro Glu Asp Val
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Pro Pro Glu Ile Leu Ser Asn Glu Arg Tyr Ala Leu Gln Lys Ala Asn
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Val Glu Glu Thr Ile Gly Arg His Val Leu Gly Ile Leu Asp Arg Ser
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Asn Met Trp Ser Lys Val Thr Glu Glu Gly Thr Glu Leu Ser Gln Arg
    1810              1815              1820
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Val Thr Gln Thr Glu Leu Met Arg Glu Ser Phe Arg Gln Lys Gln Glu
          1875   1880   . 1885
Ala Thr Glu Ser Leu Lys Cys Gln Glu Glu Leu Arg Glu Arg Leu His
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Glu Glu Ser Arg Ala Arg Glu Gln Leu Ala Val Glu Leu Ser Lys Ala
1905   1910   1915   1920
Glu Gly Val Ile Asp Gly Tyr Ala Asp Glu Lys Thr Leu Phe Glu Arg
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Gln Ile Gln Glu Lys Thr Asp Ile Ile Asp Arg Leu Glu Gln Glu Leu
          1940   1945   1950
Leu Cys Ala Ser Asn Arg Leu Gln Glu Leu Glu Ala Glu Gln Gln Gln
          1955   1960   .   1965
Ile Gln Glu Glu Arg Glu Leu Leu Ser Arg Gln Lys Glu Ala Met Lys.
          1970   1975   1980
Ala Glu Ala Gly Pro Val Glu Gln Gln Leu Leu Gln Glu Thr Glu Lys
1985   1990   1995   2000
Leu Met Lys Glu Lys Leu Glu Val Gln Cys Gln Ala Glu Lys Val Arg
          2005   2010   2015
Asp Asp Leu Gln Lys Gln Val Lys Ala Leu Glu Ile Asp Val Glu Glu
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Gln Val Ser Arg Phe Ile Glu Leu Glu Gln Glu Lys Asn Thr Glu Leu
   .   2035   2040   2045
Met Asp Leu Arg Gln Gln Asn Gln Ala Leu Glu Lys Gln Leu Glu Lys
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Met Arg Lys Phe Leu Asp Glu Gln Ala Ile Asp Arg Glu His Glu Arg
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Val Pro Arg Phe Gln Pro Ile Ser Glu His Gln Thr Arg Glu Val Glu
          2100   2105   2110
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Glu Ile Glu Lys Leu Glu Phe Arg Val Arg Glu Leu Glu Gln Ala Leu
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Leu Val Ser Ala Asp Thr Phe Gln Lys Val Glu Asp Arg Lys His Phe
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Gly Ala Val Glu Ala Lys Pro Glu Leu Ser Leu Glu Val Gln Leu Gln
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Ala Glu Arg Asp Ala Ile Asp Arg Lys Glu Lys Glu Ile Thr Asn Leu
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Glu Val Gln Gln Leu His Met Gln Leu Glu Ile Gln Lys Lys Glu Ser
2225   2230   2235   2240
Thr Thr Arg Leu Gln Glu Leu Glu Gln Glu Asn Lys Leu Phe Lys Asp
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Asp Met Glu Lys Leu Gly Leu Ala Ile Lys Glu Ser Asp Ala Met Ser
          2260   .   2265   2270
Thr Gln Asp Gln His Val Leu Phe Gly Lys Phe Ala Gln Ile Ile Gln
          2275   2280   2285
Glu Lys Glu Val Glu Ile Asp Gln Leu Asn Glu Gln Val Thr Lys Leu

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Asn Glu Leu Ile Arg Asp Leu Glu Thr Gln Ile Glu Cys Leu Met Ser
            2325                2330                2335
Asp Gln Glu Cys Val Lys Arg Asn Arg Glu Glu Glu Ile Glu Gln Leu
            2340                2345                2350
Asn Glu Val Ile Glu Lys Leu Gln Gln Glu Leu Ala Asn Ile Gly Gln
            2355                2360                2365
Lys Thr Ser Met Asn Ala His Ser Leu Ser Glu Glu Ala Asp Ser Leu
            2370                2375                2380
Lys His Gln Leu Asp Val Val Ile Ala Glu Lys Leu Ala Leu Glu Gln
2385                2390                2395                2400
Gln Val Glu Thr Ala Asn Glu Glu Met Thr Phe Met Lys Asn Val Leu
            2405                2410                2415
Lys Glu Thr Asn Phe Lys Met Asn Gln Leu Thr Gln Glu Leu Phe Ser
            2420                2425                2430
Leu Lys Arg Glu Arg Glu Ser Val Glu Lys Ile Gln Ser Ile Pro Glu
            2435                2440                2445
Asn Ser Val Asn Val Ala Ile Asp His Leu Ser Lys Asp Lys Pro Glu
            2450                2455                2460
Leu Glu Val Val Leu Thr Glu Asp Ala Leu Lys Ser Leu Glu Asn Gln
2465                2470                2475                2480
Thr Tyr Phe Lys Ser Phe Glu Glu Asn Gly Lys Gly Ser Ile Ile Asn
            2485                2490                2495
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            2500                2505                2510
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Gly Gln Phe Glu Thr Glu Met Leu Gln Lys Lys Ile Val Asn Leu Gln
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Lys Ile Val Glu Glu Lys Val Ala Ala Ala Leu Val Ser Gln Ile Gln
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            2580                2585                2590
Arg Glu Asp Glu Leu Gly Ser Asp Ile Ser Ala Leu Thr Leu Arg Ile
            2595                2600                2605
Ser Glu Leu Glu Ser Gln Val Val Glu Met His Thr Ser Leu Ile Leu
            2610                2615                2620
Glu Lys Glu Gln Val Glu Ile Ala Glu Lys Asn Val Leu Glu Lys Glu
2625                2630                2635                2640
Lys Lys Leu Leu Glu Leu Gln Lys Leu Leu Glu Gly Asn Glu Lys Lys
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            2660                2665                2670
Lys Thr Thr Thr Glu Leu Phe His Ser Asn Glu Glu Ser Gly Phe Phe
            2675                2680                2685
Asn Glu Leu Glu Ala Leu Arg Ala Glu Ser Val Ala Thr Lys Ala Glu
            2690                2695                2700
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Val Lys Glu Thr Asn Met Thr Ser Leu Gln Lys Asp Leu Ser Gln Val
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Ser Lys Lys Gln Lys Lys Lys Arg Lys Thr Ser Ser Ser Lys His Asp
         35                  40                  45
Val Ser Ala His His Asp Leu Asn Ile Asp Gln Ser Gln Cys Asn Glu
     50                  55                  60
Met Tyr Ile Asn Ser Ser Gln Arg Val Glu Ser Thr Val Ile Pro Glu
 65                  70                  75                  80
Ser Thr Ile Met Arg Thr Leu His Ser Gly Glu Ile Thr Ser His Glu
                 85                  90                  95
Gln Gly Phe Ser Val Glu Leu Glu Ser Glu Ile Ser Thr Thr Ala Asp
            100                 105                 110
Asp Cys Ser Ser Glu Val Asn Gly Cys Ser Phe Val Met Arg Thr Gly
        115                 120                 125
Lys Pro Thr Asn Leu Leu Arg Glu Glu Glu Phe Gly Val Asp Asp Ser
    130                 135                 140
Tyr Ser Glu Gln Gly Ala Gln Asp Ser Pro Thr His Leu Glu Met Met
145                 150                 155                 160
Glu Ser Glu Leu Ala Gly Lys Gln His Glu Ile Glu Glu Leu Asn Arg
                165                 170                 175
Glu Leu Glu Glu Met Arg Val Thr Tyr Gly Thr Glu Gly Leu Gln Gln
            180                 185                 190
Leu Gln Glu Phe Glu Ala Ala Ile Lys Gln Arg Asp Gly Ile Ile Thr
        195                 200                 205
Gln Leu Thr Ala Asn Leu Gln Gln Ala Arg Arg Glu Lys Asp Glu Thr
    210                 215                 220
Met Arg Glu Phe Leu Glu Leu Thr Glu Gln Ser Gln Lys Leu Gln Ile
225                 230                 235                 240
Gln Phe Gln Gln Leu Gln Ala Ser Glu Thr Leu Arg Asn Ser Thr His
            245                 250                 255
Ser Ser Thr Ala Ala Asp Leu Leu Gln Ala Lys Gln Gln Ile Leu Thr
        260                 265                 270
His Gln Gln Gln Leu Glu Glu Gln Asp His Leu Leu Glu Asp Tyr Gln
    275                 280                 285
Lys Lys Lys Glu Asp Phe Thr Met Gln Ile Ser Phe Leu Gln Glu Lys
    290                 295                 300
Ile Lys Val Tyr Glu Met Glu Gln Asp Lys Lys Val Glu Asn Ser Asn
305                 310                 315                 320
Lys Glu Glu Ile Gln Glu Lys Glu Thr Ile Ile Glu Glu Leu Asn Thr
            325                 330                 335
Lys Ile Ile Glu Glu Glu Lys Lys Thr Leu Glu Leu Lys Asp Lys Leu
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Thr Thr Ala Asp Lys Leu Leu Gly Glu Leu Gln Glu Gln Ile Val Gln
        355                 360                 365
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Gln Lys Glu Arg Gln Ser Ser Glu Glu Ile Lys Gln Leu Met Gly Thr
385                 390             395                 400
Val Glu Glu Leu Gln Lys Arg Asn His Lys Asp Ser Gln Phe Glu Thr
            405             410             415
Asp Ile Val Gln Arg Met Glu Gln Glu Thr Gln Arg Lys Leu Glu Gln
            420             425             430
Leu Arg Ala Glu Leu Asp Glu Met Tyr Gly Gln Gln Ile Val Gln Met
        435             440             445
Lys Gln Glu Leu Ile Arg Gln His Met Ala Gln Met Glu Glu Met Lys
    450             455             460
Thr Arg His Lys Gly Glu Met Glu Asn Ala Leu Arg Ser Tyr Ser Asn
465             470             475                 480
Ile Thr Val Asn Glu Asp Gln Ile Lys Leu Met Asn Val Ala Ile Asn
            485             490             495
Glu Leu Asn Ile Lys Leu Gln Asp Thr Asn Ser Gln Lys Glu Lys Leu
            500             505             510
Lys Glu Glu Leu Gly Leu Ile Leu Glu Glu Lys Cys Ala Leu Gln Arg
        515             520             525
Gln Leu Glu Asp Leu Val Glu Glu Leu Ser Phe Ser Arg Glu Gln Ile
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Gln Arg Ala Arg Gln Thr Ile Ala Glu Gln Glu Ser Lys Leu Asn Glu
545             550             555                 560
Ala His Lys Ser Leu Ser Thr Val Glu Asp Leu Lys Ala Glu Ile Val
            565             570             575
Ser Ala Ser Glu Ser Arg Lys Glu Leu Glu Leu Lys His Glu Ala Glu
            580             585             590
Val Thr Asn Tyr Lys Ile Lys Leu Glu Met Leu Glu Lys Glu Lys Asn
    595             600             605
Ala Val Leu Asp Arg Met Ala Glu Ser Gln Glu Ala Glu Leu Glu Arg
    610             615             620
Leu Arg Thr Gln Leu Leu Phe Ser His Glu Glu Glu Leu Ser Lys Leu
625             630             635                 640
Lys Glu Asp Leu Glu Ile Glu His Arg Ile Asn Ile Glu Lys Leu Lys
            645             650             655
Asp Asn Leu Gly Ile His Tyr Lys Gln Gln Ile Asp Gly Leu Gln Asn
            660             665             670
Glu Met Ser Gln Lys Ile Glu Thr Met Gln Phe Glu Lys Asp Asn Leu
        675             680             685
Ile Thr Lys Gln Asn Gln Leu Ile Leu Glu Ile Ser Lys Leu Lys Asp
    690             695             700
Leu Gln Gln Ser Leu Val Asn Ser Lys Ser Glu Glu Met Thr Leu Gln
705                 710             715                 720
Ile Asn Glu Leu Gln Lys Glu Ile Glu Ile Leu Arg Gln Glu Glu Lys
            725             730             735
Glu Lys Gly Thr Leu Glu Gln Glu Val Gln Glu Leu Gln Leu Lys Thr
            740             745             750
Glu Leu Leu Glu Lys Gln Met Lys Glu Lys Glu Asn Asp Leu Gln Glu
        755             760             765
Lys Phe Ala Gln Leu Glu Ala Glu Asn Ser Ile Leu Lys Asp Glu Lys
    770             775             780
Lys Thr Leu Glu Asp Met Leu Lys Ile His Thr Pro Val Ser Gln Glu
785             790             795                 800
Glu Arg Leu Ile Phe Leu Asp Ser Ile Lys Ser Lys Ser Lys Asp Ser
            805             810             815
Val Trp Glu Lys Glu Ile Glu Ile Leu Ile Glu Glu Asn Glu Asp Leu
            820             825             830
Lys Gln Gln Cys Ile Gln Leu Asn Glu Glu Ile Glu Lys Gln Arg Asn
```

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              835                    840                    845
Thr Phe Ser Phe Ala Glu Lys Asn Phe Glu Val Asn Tyr Gln Glu Leu
    850                    855                    860
Gln Glu Glu Tyr Ala Cys Leu Leu Lys Val Lys Asp Asp Leu Glu Asp
865                    870                    875                    880
Ser Lys Asn Lys Gln Glu Leu Glu Tyr Lys Ser Lys Leu Lys Ala Leu
                885                    890                    895
Asn Glu Glu Leu His Leu Gln Arg Ile Asn Pro Thr Thr Val Lys Met
                900                    905                    910
Lys Ser Ser Val Phe Asp Glu Asp Lys Thr Phe Val Ala Glu Thr Leu
                915                    920                    925
Glu Met Gly Glu Val Val Glu Lys Asp Thr Thr Glu Leu Met Glu Lys
    930                    935                    940
Leu Glu Val Thr Lys Arg Glu Lys Leu Glu Leu Ser Gln Arg Leu Ser
945                    950                    955                    960
Asp Leu Ser Glu Gln Leu Lys Gln Lys His Gly Glu Ile Ser Phe Leu
                965                    970                    975
Asn Glu Glu Val Lys Ser Leu Lys Gln Glu Lys Glu Gln Val Ser Leu
                980                    985                    990
Arg Cys Arg Glu Leu Glu Ile Ile Ile Asn His Asn Arg Ala Glu Asn
                995                    1000                   1005
Val Gln Ser Cys Asp Thr Gln Val Ser Ser Leu Leu Asp Gly Val Val
    1010                   1015                   1020
Thr Met Thr Ser Arg Gly Ala Glu Gly Ser Val Ser Lys Val Asn Lys
1025                   1030                   1035                   1040
Ser Phe Gly Glu Glu Ser Lys Ile Met Val Glu Asp Lys Val Ser Phe
                1045                   1050                   1055
Glu Asn Met Thr Val Gly Glu Glu Ser Lys Gln Glu Gln Leu Ile Leu
                1060                   1065                   1070
Asp His Leu Pro Ser Val Thr Lys Glu Ser Ser Leu Arg Ala Thr Gln
                1075                   1080                   1085
Pro Ser Glu Asn Asp Lys Leu Gln Lys Glu Leu Asn Val Leu Lys Ser
    1090                   1095                   1100
Glu Gln Asn Asp Leu Arg Leu Gln Met Glu Ala Gln Arg Ile Cys Leu
1105                   1110                   1115                   1120
Ser Leu Val Tyr Ser Thr His Val Asp Gln Val Arg Glu Tyr Met Glu
                1125                   1130                   1135
Asn Glu Lys Asp Lys Ala Leu Cys Ser Leu Lys Glu Glu Leu Ile Phe
                1140                   1145                   1150
Ala Gln Glu Glu Lys Ile Lys Glu Leu Gln Lys Ile His Gln Leu Glu
                1155                   1160                   1165
Leu Gln Thr Met Lys Thr Gln Glu Thr Gly Asp Glu Gly Lys Pro Leu
    1170                   1175                   1180
His Leu Leu Ile Gly Lys Leu Gln Lys Ala Val Ser Glu Glu Cys Ser
1185                   1190                   1195                   1200
Tyr Phe Leu Gln Thr Leu Cys Ser Val Leu Gly Glu Tyr Tyr Thr Pro
                1205                   1210                   1215
Ala Leu Lys Cys Glu Val Asn Ala Glu Asp Lys Glu Asn Ser Gly Asp
                1220                   1225                   1230
Tyr Ile Ser Glu Asn Glu Asp Pro Glu Leu Gln Asp Tyr Arg Tyr Glu
    1235                   1240                   1245
Val Gln Asp Phe Gln Glu Asn Met His Thr Leu Leu Asn Lys Val Thr
    1250                   1255                   1260
Glu Glu Tyr Asn Lys Leu Leu Val Leu Gln Thr Arg Leu Ser Lys Ile
1265                   1270                   1275                   1280
Trp Gly Gln Gln Thr Asp Gly Met Lys Leu Glu Phe Gly Glu Glu Asn
                1285                   1290                   1295
Leu Pro Lys Glu Glu Thr Glu Phe Leu Ser Ile His Ser Gln Met Thr
    1300                   1305                   1310
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Asn Leu Glu Asp Ile Asp Val Asn His Lys Ser Lys Leu Ser Ser Leu
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Gln Asp Leu Glu Lys Thr Lys Leu Glu Glu Gln Val Gln Glu Leu Glu
        1330   1335   1340
Ser Leu Ile Ser Ser Leu Gln Gln Gln Leu Lys Glu Thr Glu Gln Asn
1345   1350   1355   1360
Tyr Glu Ala Glu Ile His Cys Leu Gln Lys Arg Leu Gln Ala Val Ser
                1365   1370   1375
Glu Ser Thr Val Pro Pro Ser Leu Pro Val Asp Ser Val Val Ile Thr
                1380   1385   1390
Glu Ser Asp Ala Gln Arg Thr Met Tyr Pro Gly Ser Cys Val Lys Lys
                1395   1400   1405
Asn Ile Asp Gly Thr Ile Glu Phe Ser Gly Glu Phe Gly Val Lys Glu
        1410   1415   1420
Glu Thr Asn Ile Val Lys Leu Leu Glu Lys Gln Tyr Gln Glu Gln Leu
1425   1430   1435   1440
Glu Glu Glu Val Ala Lys Val Ile Val Ser Met Ser Ile Ala Phe Ala
                1445   1450   1455
Gln Gln Thr Glu Leu Ser Arg Ile Ser Gly Gly Lys Glu Asn Thr Ala
                1460   1465   1470
Ser Ser Lys Gln Ala His Ala Val Cys Gln Gln Glu Gln His Tyr Phe
                1475   1480   1485
Asn Glu Met Lys Leu Ser Gln Asp Gln Ile Gly Phe Gln Thr Phe Glu
        1490   1495   1500
Thr Val Asp Val Lys Phe Lys Glu Glu Phe Lys Pro Leu Ser Lys Glu
1505   1510   1515   1520
Leu Gly Glu His Gly Lys Glu Ile Leu Leu Ser Asn Ser Asp Pro His
                1525   1530   1535
Asp Ile Pro Glu Ser Lys Asp Cys Val Leu Thr Ile Ser Glu Glu Met
        1540   1545   1550
Phe Ser Lys Asp Lys Thr Phe Ile Val Arg Gln Ser Ile His Asp Glu
        1555   1560   1565
Ile Ser Val Ser Ser Met Asp Ala Ser Arg Gln Leu Met Leu Asn Glu
        1570   1575   1580
Glu Gln Leu Glu Asp Met Arg Gln Glu Leu Val Arg Gln Tyr Gln Glu
1585   1590   1595   1600
His Gln Gln Ala Thr Glu Leu Leu Arg Gln Ala His Met Arg Gln Met
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Glu Arg Gln Arg Glu Asp Gln Glu Gln Leu Gln Glu Glu Ile Lys Arg
        1620   1625   1630
Leu Asn Arg Gln Leu Ala Gln Arg Ser Ser Ile Asp Asn Glu Asn Leu
        1635   1640   1645
Val Ser Glu Arg Glu Arg Val Leu Leu Glu Glu Leu Glu Ala Leu Lys
        1650   1655   1660
Gln Leu Ser Leu Ala Gly Arg Glu Lys Leu Cys Cys Glu Leu Arg Asn
1665   1670   1675   1680
Ser Ser Thr Gln Thr Gln Asn Gly Asn Glu Asn Gln Gly Glu Val Glu
                1685   1690   1695
Glu Gln Thr Phe Lys Glu Lys Glu Leu Asp Arg Lys Pro Glu Asp Val
                1700   1705   1710
Pro Pro Glu Ile Leu Ser Asn Glu Arg Tyr Ala Leu Gln Lys Ala Asn
        1715   1720   1725
Asn Arg Leu Leu Lys Ile Leu Leu Glu Val Val Lys Thr Thr Ala Ala
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Val Glu Glu Thr Ile Gly Arg His Val Leu Gly Ile Leu Asp Arg Ser
1745   1750   1755   1760
Ser Lys Ser Gln Ser Ser Ala Ser Leu Ile Trp Arg Ser Glu Ala Glu
                1765   1770   1775
Ala Ser Val Lys Ser Cys Val His Glu Glu His Thr Arg Val Thr Asp

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       Asn Met Trp Ser Lys Val Thr Glu Glu Gly Thr Glu Leu Ser Gln Arg
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       Leu Val Arg Ser Gly Phe Ala Gly Thr Glu Ile Asp Pro Glu Asn Glu
       1825                 1830                1835                1840
       Glu Leu Met Leu Asn Ile Ser Ser Arg Leu Gln Ala Ala Val Glu Lys
                 1845                 1850                1855
       Leu Leu Glu Ala Ile Ser Glu Thr Ser Ser Gln Leu Glu His Ala Lys
              1860                 1865                1870
       Val Thr Gln Thr Glu Leu Met Arg Glu Ser Phe Arg Gln Lys Gln Glu
             1875                 1880                1885
       Ala Thr Glu Ser Leu Lys Cys Gln Glu Glu Leu Arg Glu Arg Leu His
          1890                 1895                1900
       Glu Glu Ser Arg Ala Arg Glu Gln Leu Ala Val Glu Leu Ser Lys Ala
       1905                 1910                1915                1920
       Glu Gly Val Ile Asp Gly Tyr Ala Asp Glu Lys Thr Leu Phe Glu Arg
                 1925                 1930                1935
       Gln Ile Gln Glu Lys Thr Asp Ile Ile Asp Arg Leu Glu Gln Glu Leu
              1940                 1945                1950
       Leu Cys Ala Ser Asn Arg Leu Gln Glu Leu Glu Ala Glu Gln Gln Gln
              1955                 1960                1965
       Ile Gln Glu Glu Arg Glu Leu Leu Ser Arg Gln Lys Glu Ala Met Lys
          1970                 1975                1980
       Ala Glu Ala Gly Pro Val Glu Gln Gln Leu Leu Gln Glu Thr Glu Lys
       1985                 1990                1995                2000
       Leu Met Lys Glu Lys Leu Glu Val Gln Cys Gln Ala Glu Lys Val Arg
                 2005                 2010                2015
       Asp Asp Leu Gln Lys Gln Val Lys Ala Leu Glu Ile Asp Val Glu Glu
              2020                 2025                2030
       Gln Val Ser Arg Phe Ile Glu Leu Glu Gln Glu Lys Asn Thr Glu Leu
              2035                 2040                2045
       Met Asp Leu Arg Gln Gln Asn Gln Ala Leu Glu Lys Gln Leu Glu Lys
          2050                 2055                2060
       Met Arg Lys Phe Leu Asp Glu Gln Ala Ile Asp Arg Glu His Glu Arg
       2065                 2070                2075                2080
       Asp Val Phe Gln Gln Glu Ile Gln Lys Leu Glu Gln Gln Leu Lys Val
                 2085                 2090                2095
       Val Pro Arg Phe Gln Pro Ile Ser Glu His Gln Thr Arg Glu Val Glu
                 2100                 2105                2110
       Gln Leu Ala Asn His Leu Lys Glu Lys Thr Asp Lys Cys Ser Glu Leu
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       Leu Leu Ser Lys Glu Gln Leu Gln Arg Asp Ile Gln Glu Arg Asn Glu
          2130                 2135                2140
       Glu Ile Glu Lys Leu Glu Phe Arg Val Arg Glu Leu Glu Gln Ala Leu
       2145                 2150                2155                2160
       Leu Val Glu Asp Arg Lys His Phe Gly Ala Val Glu Ala Lys Pro Glu
                 2165                 2170                2175
       Leu Ser Leu Glu Val Gln Leu Gln Ala Glu Arg Asp Ala Ile Asp Arg
              2180                 2185                2190
       Lys Glu Lys Glu Ile Thr Asn Leu Glu Glu Gln Leu Glu Gln Phe Arg
              2195                 2200                2205
       Glu Glu Leu Glu Asn Lys Asn Glu Glu Val Gln Gln Leu His Met Gln
          2210                 2215                2220
       Leu Glu Ile Gln Lys Lys Glu Ser Thr Thr Arg Leu Gln Glu Leu Glu
       2225                 2230                2235                2240
       Gln Glu Asn Lys Leu Phe Lys Asp Asp Met Glu Lys Leu Gly Leu Ala
                 2245                 2250                2255
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Gly Lys Phe Ala Gln Ile Ile Gln Glu Lys Glu Val Glu Ile Asp Gln
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Leu Asn Glu Gln Val Thr Lys Leu Gln Gln Gln Leu Lys Ile Thr Thr
        2290                2295                2300
Asp Asn Lys Val Ile Glu Glu Lys Asn Glu Leu Ile Arg Asp Leu Glu
2305                2310                2315                2320
Thr Gln Ile Glu Cys Leu Met Ser Asp Gln Glu Cys Val Lys Arg Asn
        2325                2330                2335
Arg Glu Glu Glu Ile Glu Gln Leu Asn Glu Val Ile Glu Lys Leu Gln
        2340                2345                2350
Gln Glu Leu Ala Asn Ile Gly Gln Lys Thr Ser Met Asn Ala His Ser
        2355                2360                2365
Leu Ser Glu Glu Ala Asp Ser Leu Lys His Gln Leu Asp Val Val Ile
        2370                2375                2380
Ala Glu Lys Leu Ala Leu Glu Gln Gln Val Glu Thr Ala Asn Glu Glu
2385                2390                2395                2400
Met Thr Phe Met Lys Asn Val Leu Lys Glu Thr Asn Phe Lys Met Asn
        2405                2410                2415
Gln Leu Thr Gln Glu Leu Phe Ser Leu Lys Arg Glu Arg Glu Ser Val
        2420                2425                2430
Glu Lys Ile Gln Ser Ile Pro Glu Asn Ser Val Asn Val Ala Ile Asp
        2435                2440                2445
His Leu Ser Lys Asp Lys Pro Glu Leu Glu Val Val Leu Thr Glu Asp
        2450                2455                2460
Ala Leu Lys Ser Leu Glu Asn Gln Thr Tyr Phe Lys Ser Phe Glu Glu
2465                2470                2475                2480
Asn Gly Lys Gly Ser Ile Ile Asn Leu Glu Thr Arg Leu Leu Gln Leu
        2485                2490                2495
Glu Ser Thr Val Ser Ala Lys Asp Leu Glu Leu Thr Gln Cys Tyr Lys
        2500                2505                2510
Gln Ile Lys Asp Met Gln Glu Gln Gly Gln Phe Glu Thr Glu Met Leu
        2515                2520                2525
Gln Lys Lys Ile Val Asn Leu Gln Lys Ile Val Glu Glu Lys Val Ala
        2530                2535                2540
Ala Ala Leu Val Ser Gln Ile Gln Leu Glu Ala Val Gln Glu Tyr Ala
2545                2550                2555                2560
Lys Phe Cys Gln Asp Asn Gln Thr Ile Ser Ser Glu Pro Glu Arg Thr
        2565                2570                2575
Asn Ile Gln Asn Leu Asn Gln Leu Arg Glu Asp Glu Leu Gly Ser Asp
        2580                2585                2590
Ile Ser Ala Leu Thr Leu Arg Ile Ser Glu Leu Glu Ser Gln Val Val
        2595                2600                2605
Glu Met His Thr Ser Leu Ile Leu Glu Lys Glu Gln Val Glu Ile Ala
        2610                2615                2620
Glu Lys Asn Val Leu Glu Lys Glu Lys Leu Leu Glu Leu Gln Lys
2625                2630                2635                2640
Leu Leu Glu Gly Asn Glu Lys Lys Gln Arg Glu Lys Glu Lys Lys Arg
        2645                2650                2655
Ser Pro Gln Asp Val Glu Val Leu Lys Thr Thr Thr Glu Leu Phe His
        2660                2665                2670
Ser Asn Glu Ser Gly Phe Phe Asn Glu Leu Glu Ala Leu Arg Ala
        2675                2680                2685
Glu Ser Val Ala Thr Lys Ala Glu Leu Ala Ser Tyr Lys Glu Lys Ala
        2690                2695                2700
Glu Lys Leu Gln Glu Glu Leu Leu Val Lys Glu Thr Asn Met Thr Ser
2705                2710                2715                2720
Leu Gln Lys Asp Leu Ser Gln Val Arg Asp His Leu Ala Glu Ala Lys
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Glu Lys Leu Ser Ile Leu Glu Lys Glu Asp Glu Thr Glu Val Gln Glu
          2740                    2745                    2750
Ser Lys Lys Ala Cys Met Phe Glu Pro Leu Pro Ile Lys Leu Ser Lys
          2755                    2760                    2765
Ser Ile Ala Ser Gln Thr Asp Gly Thr Leu Lys Ile Ser Ser Ser Asn
      2770                    2775                    2780
Gln Thr Pro Gln Ile Leu Val Lys Asn Ala Gly Ile Gln Ile Asn Leu
2785                    2790                    2795                    2800
Gln Ser Glu Cys Ser Ser Glu Glu Val Thr Glu Ile Ile Ser Gln Phe
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Thr Glu Lys Ile Glu Lys Met Gln Glu Leu His Ala Ala Glu Ile Leu
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Asp Met Glu Ser Arg His Ile Ser Glu Thr Glu Thr Leu Lys Arg Glu
          2835                    2840                    2845
His Tyr Val Ala Val Gln Leu Leu Lys Glu Glu Cys Gly Thr Leu Lys
      2850                    2855                    2860
Ala Val Ile Gln Cys Leu Arg Ser Lys Glu Gly Ser Ser Ile Pro Glu
2865                    2870                    2875                    2880
Leu Ala His Ser Asp Ala Tyr Gln Thr Arg Glu Ile Cys Ser Ser Asp
              2885                    2890                    2895
Ser Gly Ser Asp Trp Gly Gln Gly Ile Tyr Leu Thr His Ser Gln Gly
          2900                    2905                    2910
Phe Asp Ile Ala Ser Glu Gly Arg Gly Glu Glu Ser Glu Ser Ala Thr
      2915                    2920                    2925
Asp Ser Phe Pro Lys Lys Ile Lys Gly Leu Leu Arg Ala Val His Asn
      2930                    2935                    2940
Glu Gly Met Gln Val Leu Ser Leu Thr Glu Ser Pro Tyr Ser Asp Gly
2945                    2950                    2955                    2960
Glu Asp His Ser Ile Gln Gln Val Ser Glu Pro Trp Leu Glu Glu Arg
              2965                    2970                    2975
Lys Ala Tyr Ile Asn Thr Ile Ser Ser Leu Lys Asp Leu Ile Thr Lys
          2980                    2985                    2990
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agactactga gacaaaatgc tgagctgaca gggcatatca gtcaactgac tgaagagaag 11040
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```

29

```
gcagaaaaga ctgactcatt ttatcattct tctggtgggc tggagttata tggagaacca 11700
agacatacta cgtatcgctc aagatcagat ctggactata ttaggtcccc tttaccattt 11760
cagaataggt acccaggcac tccagctgat ttcaatcctg gttctttagc atgttctcag 11820
cttcagaatt acgatcctga cagagcccta acagattata tcactcggct agaggcactg 11880
caaagacgac ttggaactat acagtcaggt gctctgagtt taaccacatc ttggcagcac 11940
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tcccantcct cccaaaaatg atttcctact atgttcattc agcggactga tgacacaaaa 12180
tgcacaatga gcaccagtgt gcaaggtact ctgagtttac agagcctaac tggagaacgt 12240
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agcagagcga gaccctgtct caaagaaaaa aaaaaaa                          12337
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<211> 3925
<212> PRT
<213> Homo sapiens

<400> 6

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 1               5                  10                  15
Leu Ala Gln Phe Arg Gln Arg Lys Ala Gln Ser Asp Gly Gln Ser Pro
            20                  25                  30
Ser Lys Lys Gln Lys Lys Lys Arg Lys Thr Ser Ser Ser Lys His Asp
        35                  40                  45
Val Ser Ala His His Asp Leu Asn Ile Asp Gln Ser Gln Cys Asn Glu
    50                  55                  60
Met Tyr Ile Asn Ser Ser Gln Arg Val Glu Ser Thr Val Ile Pro Glu
65                  70                  75                  80
Ser Thr Ile Met Arg Thr Leu His Ser Gly Glu Ile Thr Ser His Glu
                85                  90                  95
Gln Gly Phe Ser Val Glu Leu Glu Ser Glu Ile Ser Thr Thr Ala Asp
            100                 105                 110
Asp Cys Ser Ser Glu Val Asn Gly Cys Ser Phe Val Met Arg Thr Gly
        115                 120                 125
Lys Pro Thr Asn Leu Leu Arg Glu Glu Glu Phe Gly Val Asp Asp Ser
    130                 135                 140
Tyr Ser Glu Gln Gly Ala Gln Asp Ser Pro Thr His Leu Glu Met Met
145                 150                 155                 160
Glu Ser Glu Leu Ala Gly Lys Gln His Glu Ile Glu Glu Leu Asn Arg
                165                 170                 175
Glu Leu Glu Glu Met Arg Val Thr Tyr Gly Thr Glu Gly Leu Gln Gln
            180                 185                 190
Leu Gln Glu Phe Glu Ala Ala Ile Lys Gln Arg Asp Gly Ile Ile Thr
        195                 200                 205
Gln Leu Thr Ala Asn Leu Gln Gln Ala Arg Arg Glu Lys Asp Glu Thr
    210                 215                 220
Met Arg Glu Phe Leu Glu Leu Thr Glu Gln Ser Gln Lys Leu Gln Ile
225                 230                 235                 240
Gln Phe Gln Gln Leu Gln Ala Ser Glu Thr Leu Arg Asn Ser Thr His
            245                 250                 255
Ser Ser Thr Ala Ala Asp Leu Leu Gln Ala Lys Gln Gln Ile Leu Thr
        260                 265                 270
His Gln Gln Gln Leu Glu Glu Gln Asp His Leu Leu Glu Asp Tyr Gln
    275                 280                 285
Lys Lys Lys Glu Asp Phe Thr Met Gln Ile Ser Phe Leu Gln Glu Lys
    290                 295                 300
Ile Lys Val Tyr Glu Met Glu Gln Asp Lys Lys Val Glu Asn Ser Asn
305                 310                 315                 320
```

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Lys Glu Glu Ile Gln Glu Lys Glu Thr Ile Ile Glu Glu Leu Asn Thr
            325                   330                   335
Lys Ile Ile Glu Glu Glu Lys Lys Thr Leu Glu Leu Lys Asp Lys Leu
            340                   345                   350
Thr Thr Ala Asp Lys Leu Leu Gly Glu Leu Gln Glu Gln Ile Val Gln
            355                   360                   365
Lys Asn Gln Glu Ile Lys Asn Met Lys Leu Glu Leu Thr Asn Ser Lys
    370                   375                   380
Gln Lys Glu Arg Gln Ser Ser Glu Glu Ile Lys Gln Leu Met Gly Thr
385                   390                   395                   400
Val Glu Glu Leu Gln Lys Arg Asn His Lys Asp Ser Gln Phe Glu Thr
            405                   410                   415
Asp Ile Val Gln Arg Met Glu Gln Glu Thr Gln Arg Lys Leu Glu Gln
            420                   425                   430
Leu Arg Ala Glu Leu Asp Glu Met Tyr Gly Gln Gln Ile Val Gln Met
            435                   440                   445
Lys Gln Glu Leu Ile Arg Gln His Met Ala Gln Met Glu Glu Met Lys
    450                   455                   460
Thr Arg His Lys Gly Glu Met Glu Asn Ala Leu Arg Ser Tyr Ser Asn
465                   470                   475                   480
Ile Thr Val Asn Glu Asp Gln Ile Lys Leu Met Asn Val Ala Ile Asn
            485                   490                   495
Glu Leu Asn Ile Lys Leu Gln Asp Thr Asn Ser Gln Lys Glu Lys Leu
            500                   505                   510
Lys Glu Glu Leu Gly Leu Ile Leu Glu Glu Lys Cys Ala Leu Gln Arg
            515                   520                   525
Gln Leu Glu Asp Leu Val Glu Glu Leu Ser Phe Ser Arg Glu Gln Ile
    530                   535                   540
Gln Arg Ala Arg Gln Thr Ile Ala Glu Gln Glu Ser Lys Leu Asn Glu
545                   550                   555                   560
Ala His Lys Ser Leu Ser Thr Val Glu Asp Leu Lys Ala Glu Ile Val
            565                   570                   575
Ser Ala Ser Glu Ser Arg Lys Glu Leu Glu Leu Lys His Glu Ala Glu
            580                   585                   590
Val Thr Asn Tyr Lys Ile Lys Leu Glu Met Leu Glu Lys Glu Lys Asn
            595                   600                   605
Ala Val Leu Asp Arg Met Ala Glu Ser Gln Glu Ala Glu Leu Glu Arg
    610                   615                   620
Leu Arg Thr Gln Leu Leu Phe Ser His Glu Glu Glu Leu Ser Lys Leu
625                   630                   635                   640
Lys Glu Asp Leu Glu Ile Glu His Arg Ile Asn Ile Glu Lys Leu Lys
            645                   650                   655
Asp Asn Leu Gly Ile His Tyr Lys Gln Gln Ile Asp Gly Leu Gln Asn
            660                   665                   670
Glu Met Ser Gln Lys Ile Glu Thr Met Gln Phe Glu Lys Asp Asn Leu
    675                   680                   685
Ile Thr Lys Gln Asn Gln Leu Ile Leu Glu Ile Ser Lys Leu Lys Asp
    690                   695                   700
Leu Gln Gln Ser Leu Val Asn Ser Lys Ser Glu Met Thr Leu Gln
705                   710                   715                   720
Ile Asn Glu Leu Gln Lys Glu Ile Glu Ile Leu Arg Gln Glu Glu Lys
            725                   730                   735
Glu Lys Gly Thr Leu Glu Gln Glu Val Gln Glu Leu Gln Leu Lys Thr
            740                   745                   750
Glu Leu Leu Glu Lys Gln Met Lys Glu Lys Glu Asn Asp Leu Gln Glu
            755                   760                   765
Lys Phe Ala Gln Leu Glu Ala Glu Asn Ser Ile Leu Lys Asp Glu Lys
    770                   775                   780
Lys Thr Leu Glu Asp Met Leu Lys Ile His Thr Pro Val Ser Gln Glu
```

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785                     790                     795                     800
Glu Arg Leu Ile Phe Leu Asp Ser Ile Lys Ser Lys Ser Lys Asp Ser
                805                     810                     815
Val Trp Glu Lys Glu Ile Glu Ile Leu Ile Glu Glu Asn Glu Asp Leu
                820                     825                     830
Lys Gln Gln Cys Ile Gln Leu Asn Glu Glu Ile Glu Lys Gln Arg Asn
            835                     840                     845
Thr Phe Ser Phe Ala Glu Lys Asn Phe Glu Val Asn Tyr Gln Glu Leu
        850                     855                     860
Gln Glu Glu Tyr Ala Cys Leu Leu Lys Val Lys Asp Asp Leu Glu Asp
865                     870                     875                     880
Ser Lys Asn Lys Gln Glu Leu Glu Tyr Lys Ser Lys Leu Lys Ala Leu
                885                     890                     895
Asn Glu Glu Leu His Leu Gln Arg Ile Asn Pro Thr Thr Val Lys Met
            900                     905                     910
Lys Ser Ser Val Phe Asp Glu Asp Lys Thr Phe Val Ala Glu Thr Leu
            915                     920                     925
Glu Met Gly Glu Val Val Glu Lys Asp Thr Thr Glu Leu Met Glu Lys
        930                     935                     940
Leu Glu Val Thr Lys Arg Glu Lys Leu Glu Leu Ser Gln Arg Leu Ser
945                     950                     955                     960
Asp Leu Ser Glu Gln Leu Lys Gln Lys His Gly Glu Ile Ser Phe Leu
            965                     970                     975
Asn Glu Glu Val Lys Ser Leu Lys Gln Glu Lys Glu Gln Val Ser Leu
            980                     985                     990
Arg Cys Arg Glu Leu Glu Ile Ile Ile Asn His Asn Arg Ala Glu Asn
        995                     1000                    1005
Val Gln Ser Cys Asp Thr Gln Val Ser Ser Leu Leu Asp Gly Val Val
    1010                    1015                    1020
Thr Met Thr Ser Arg Gly Ala Glu Gly Ser Val Ser Lys Val Asn Lys
1025                    1030                    1035                    1040
Ser Phe Gly Glu Glu Ser Lys Ile Met Val Glu Asp Lys Val Ser Phe
                1045                    1050                    1055
Glu Asn Met Thr Val Gly Glu Glu Ser Lys Gln Glu Gln Leu Ile Leu
            1060                    1065                    1070
Asp His Leu Pro Ser Val Thr Lys Glu Ser Ser Leu Arg Ala Thr Gln
            1075                    1080                    1085
Pro Ser Glu Asn Asp Lys Leu Gln Lys Glu Leu Asn Val Leu Lys Ser
    1090                    1095                    1100
Glu Gln Asn Asp Leu Arg Leu Gln Met Glu Ala Gln Arg Ile Cys Leu
1105                    1110                    1115                    1120
Ser Leu Val Tyr Ser Thr His Val Asp Gln Val Arg Glu Tyr Met Glu
                1125                    1130                    1135
Asn Glu Lys Asp Lys Ala Leu Cys Ser Leu Lys Glu Glu Leu Ile Phe
            1140                    1145                    1150
Ala Gln Glu Glu Lys Ile Lys Glu Leu Gln Lys Ile His Gln Leu Glu
            1155                    1160                    1165
Leu Gln Thr Met Lys Thr Gln Glu Thr Gly Asp Glu Gly Lys Pro Leu
    1170                    1175                    1180
His Leu Leu Ile Gly Lys Leu Gln Lys Ala Val Ser Glu Glu Cys Ser
1185                    1190                    1195                    1200
Tyr Phe Leu Gln Thr Leu Cys Ser Val Leu Gly Glu Tyr Tyr Thr Pro
                1205                    1210                    1215
Ala Leu Lys Cys Glu Val Asn Ala Glu Asp Lys Glu Asn Ser Gly Asp
            1220                    1225                    1230
Tyr Ile Ser Glu Asn Glu Asp Pro Glu Leu Gln Asp Tyr Arg Tyr Glu
            1235                    1240                    1245
Val Gln Asp Phe Gln Glu Asn Met His Thr Leu Leu Asn Lys Val Thr
    1250                    1255                    1260
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Glu Glu Tyr Asn Lys Leu Leu Val Leu Gln Thr Arg Leu Ser Lys Ile
1265                1270                1275                1280
Trp Gly Gln Gln Thr Asp Gly Met Lys Leu Glu Phe Gly Glu Glu Asn
                1285                1290                1295
Leu Pro Lys Glu Glu Thr Glu Phe Leu Ser Ile His Ser Gln Met Thr
                1300                1305                1310
Asn Leu Glu Asp Ile Asp Val Asn His Lys Ser Lys Leu Ser Ser Leu
                1315                1320                1325
Gln Asp Leu Glu Lys Thr Lys Leu Glu Glu Gln Val Gln Glu Leu Glu
                1330                1335                1340
Ser Leu Ile Ser Ser Leu Gln Gln Gln Leu Lys Glu Thr Glu Gln Asn
1345                1350                1355                1360
Tyr Glu Ala Glu Ile His Cys Leu Gln Lys Arg Leu Gln Ala Val Ser
                1365                1370                1375
Glu Ser Thr Val Pro Pro Ser Leu Pro Val Asp Ser Val Val Ile Thr
                1380                1385                1390
Glu Ser Asp Ala Gln Arg Thr Met Tyr Pro Gly Ser Cys Val Lys Lys
                1395                1400                1405
Asn Ile Asp Gly Thr Ile Glu Phe Ser Gly Glu Phe Gly Val Lys Glu
                1410                1415                1420
Glu Thr Asn Ile Val Lys Leu Leu Glu Lys Gln Tyr Gln Glu Gln Leu
1425                1430                1435                1440
Glu Glu Glu Val Ala Lys Val Ile Val Ser Met Ser Ile Ala Phe Ala
                1445                1450                1455
Gln Gln Thr Glu Leu Ser Arg Ile Ser Gly Gly Lys Glu Asn Thr Ala
                1460                1465                1470
Ser Ser Lys Gln Ala His Ala Val Cys Gln Gln Glu Gln His Tyr Phe
                1475                1480                1485
Asn Glu Met Lys Leu Ser Gln Asp Gln Ile Gly Phe Gln Thr Phe Glu
                1490                1495                1500
Thr Val Asp Val Lys Phe Lys Glu Glu Phe Lys Pro Leu Ser Lys Glu
1505                1510                1515                1520
Leu Gly Glu His Gly Lys Glu Ile Leu Leu Ser Asn Ser Asp Pro His
                1525                1530                1535
Asp Ile Pro Glu Ser Lys Asp Cys Val Leu Thr Ile Ser Glu Glu Met
                1540                1545                1550
Phe Ser Lys Asp Lys Thr Phe Ile Val Arg Gln Ser Ile His Asp Glu
                1555                1560                1565
Ile Ser Val Ser Ser Met Asp Ala Ser Arg Gln Leu Met Leu Asn Glu
1570                1575                1580
Glu Gln Leu Glu Asp Met Arg Gln Glu Leu Val Arg Gln Tyr Gln Glu
1585                1590                1595                1600
His Gln Gln Ala Thr Glu Leu Leu Arg Gln Ala His Met Arg Gln Met
                1605                1610                1615
Glu Arg Gln Arg Glu Asp Gln Glu Gln Leu Gln Glu Glu Ile Lys Arg
                1620                1625                1630
Leu Asn Arg Gln Leu Ala Gln Arg Ser Ser Ile Asp Asn Glu Asn Leu
                1635                1640                1645
Val Ser Glu Arg Glu Arg Val Leu Leu Glu Glu Leu Glu Ala Leu Lys
                1650                1655                1660
Gln Leu Ser Leu Ala Gly Arg Glu Lys Leu Cys Cys Glu Leu Arg Asn
1665                1670                1675                1680
Ser Ser Thr Gln Thr Gln Asn Gly Asn Glu Asn Gln Gly Glu Val Glu
                1685                1690                1695
Glu Gln Thr Phe Lys Glu Lys Glu Leu Asp Arg Lys Pro Glu Asp Val
                1700                1705                1710
Pro Pro Glu Ile Leu Ser Asn Glu Arg Tyr Ala Leu Gln Lys Ala Asn
                1715                1720                1725
Asn Arg Leu Leu Lys Ile Leu Leu Glu Val Val Lys Thr Thr Ala Ala
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      1730                    1735                    1740
Val Glu Glu Thr Ile Gly Arg His Val Leu Gly Ile Leu Asp Arg Ser
1745                    1750                    1755                    1760
Ser Lys Ser Gln Ser Ser Ala Ser Leu Ile Trp Arg Ser Glu Ala Glu
                  1765                    1770                    1775
Ala Ser Val Lys Ser Cys Val His Glu Glu His Thr Arg Val Thr Asp
                  1780                    1785                    1790
Glu Ser Ile Pro Ser Tyr Ser Gly Ser Asp Met Pro Arg Asn Asp Ile
          1795                    1800                    1805
Asn Met Trp Ser Lys Val Thr Glu Glu Gly Thr Glu Leu Ser Gln Arg
          1810                    1815                    1820
Leu Val Arg Ser Gly Phe Ala Gly Thr Glu Ile Asp Pro Glu Asn Glu
1825                    1830                    1835                    1840
Glu Leu Met Leu Asn Ile Ser Ser Arg Leu Gln Ala Ala Val Glu Lys
                  1845                    1850                    1855
Leu Leu Glu Ala Ile Ser Glu Thr Ser Ser Gln Leu Glu His Ala Lys
                  1860                    1865                    1870
Val Thr Gln Thr Glu Leu Met Arg Glu Ser Phe Arg Gln Lys Gln Glu
          1875                    1880                    1885
Ala Thr Glu Ser Leu Lys Cys Gln Glu Glu Leu Arg Glu Arg Leu His
          1890                    1895                    1900
Glu Glu Ser Arg Ala Arg Glu Gln Leu Ala Val Glu Leu Ser Lys Ala
1905                    1910                    1915                    1920
Glu Gly Val Ile Asp Gly Tyr Ala Asp Glu Lys Thr Leu Phe Glu Arg
                  1925                    1930                    1935
Gln Ile Gln Glu Lys Thr Asp Ile Ile Asp Arg Leu Glu Gln Glu Leu
          1940                    1945                    1950
Leu Cys Ala Ser Asn Arg Leu Gln Glu Leu Glu Ala Glu Gln Gln Gln
          1955                    1960                    1965
Ile Gln Glu Glu Arg Glu Leu Leu Ser Arg Gln Lys Glu Ala Met Lys
          1970                    1975                    1980
Ala Glu Ala Gly Pro Val Glu Gln Gln Leu Leu Gln Glu Thr Glu Lys
1985                    1990                    1995                    2000
Leu Met Lys Glu Lys Leu Glu Val Gln Cys Gln Ala Glu Lys Val Arg
                  2005                    2010                    2015
Asp Asp Leu Gln Lys Gln Val Lys Ala Leu Glu Ile Asp Val Glu Glu
          2020                    2025                    2030
Gln Val Ser Arg Phe Ile Glu Leu Glu Gln Glu Lys Asn Thr Glu Leu
          2035                    2040                    2045
Met Asp Leu Arg Gln Gln Asn Gln Ala Leu Glu Lys Gln Leu Glu Lys
          2050                    2055                    2060
Met Arg Lys Phe Leu Asp Glu Gln Ala Ile Asp Arg Glu His Glu Arg
2065                    2070                    2075                    2080
Asp Val Phe Gln Gln Glu Ile Gln Lys Leu Glu Gln Gln Leu Lys Val
                  2085                    2090                    2095
Val Pro Arg Phe Gln Pro Ile Ser Glu His Gln Thr Arg Glu Val Glu
          2100                    2105                    2110
Gln Leu Ala Asn His Leu Lys Glu Lys Thr Asp Lys Cys Ser Glu Leu
          2115                    2120                    2125
Leu Leu Ser Lys Glu Gln Leu Gln Arg Asp Ile Gln Glu Arg Asn Glu
          2130                    2135                    2140
Glu Ile Glu Lys Leu Glu Phe Arg Val Arg Glu Leu Glu Gln Ala Leu
2145                    2150                    2155                    2160
Leu Val Ser Ala Asp Thr Phe Gln Lys Val Glu Asp Arg Lys His Phe
                  2165                    2170                    2175
Gly Ala Val Glu Ala Lys Pro Glu Leu Ser Leu Glu Val Gln Leu Gln
          2180                    2185                    2190
Ala Glu Arg Asp Ala Ile Asp Arg Lys Glu Lys Glu Ile Thr Asn Leu
          2195                    2200                    2205
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Glu Glu Gln Leu Glu Gln Phe Arg Glu Glu Leu Glu Asn Lys Asn Glu
    2210   2215   2220
Glu Val Gln Gln Leu His Met Gln Leu Glu Ile Gln Lys Lys Glu Ser
2225   2230   2235   2240
Thr Thr Arg Leu Gln Glu Leu Glu Gln Glu Asn Lys Leu Phe Lys Asp
            2245   2250   2255
Asp Met Glu Lys Leu Gly Leu Ala Ile Lys Glu Ser Asp Ala Met Ser
        2260   2265   2270
Thr Gln Asp Gln His Val Leu Phe Gly Lys Phe Ala Gln Ile Ile Gln
        2275   2280   2285
Glu Lys Glu Val Glu Ile Asp Gln Leu Asn Glu Gln Val Thr Lys Leu
        2290   2295   2300
Gln Gln Gln Leu Lys Ile Thr Thr Asp Asn Lys Val Ile Glu Glu Lys
2305   2310   2315   2320
Asn Glu Leu Ile Arg Asp Leu Glu Thr Gln Ile Glu Cys Leu Met Ser
            2325   2330   2335
Asp Gln Glu Cys Val Lys Arg Asn Arg Glu Glu Glu Ile Glu Gln Leu
        2340   2345   2350
Asn Glu Val Ile Glu Lys Leu Gln Gln Glu Leu Ala Asn Ile Gly Gln
        2355   2360   2365
Lys Thr Ser Met Asn Ala His Ser Leu Ser Glu Glu Ala Asp Ser Leu
    2370   .   2375   2380   .
Lys His Gln Leu Asp Val Val Ile Ala Glu Lys Leu Ala Leu Glu Gln
2385   2390   2395   2400
Gln Val Glu Thr Ala Asn Glu Glu Met Thr Phe Met Lys Asn Val Leu
            2405   2410   2415
Lys Glu Thr Asn Phe Lys Met Asn Gln Leu Thr Gln Glu Leu Phe Ser
            2420   2425   2430
Leu Lys Arg Glu Arg Glu Ser Val Glu Lys Ile Gln Ser Ile Pro Glu
        2435   2440   2445
Asn Ser Val Asn Val Ala Ile Asp His Leu Ser Lys Asp Lys Pro Glu
        2450   2455   2460
Leu Glu Val Val Leu Thr Glu Asp Ala Leu Lys Ser Leu Glu Asn Gln
2465   2470   2475   2480
Thr Tyr Phe Lys Ser Phe Glu Glu Asn Gly Lys Gly Ser Ile Ile Asn
            2485   2490   2495
Leu Glu Thr Arg Leu Leu Gln Leu Glu Ser Thr Val Ser Ala Lys Asp
        2500   2505   2510
Leu Glu Leu Thr Gln Cys Tyr Lys Gln Ile Lys Asp Met Gln Glu Gln
        2515   2520   2525
Gly Gln Phe Glu Thr Glu Met Leu Gln Lys Lys Ile Val Asn Leu Gln
    2530   2535   2540
Lys Ile Val Glu Glu Lys Val Ala Ala Ala Leu Val Ser Gln Ile Gln
2545   2550   2555   2560
Leu Glu Ala Val Gln Glu Tyr Ala Lys Phe Cys Gln Asp Asn Gln Thr
        2565   2570   2575
Ile Ser Ser Glu Pro Glu Arg Thr Asn Ile Gln Asn Leu Asn Gln Leu
        2580   2585   2590
Arg Glu Asp Glu Leu Gly Ser Asp Ile Ser Ala Leu Thr Leu Arg Ile
        2595   2600   .   2605
Ser Glu Leu Glu Ser Gln Val Val Glu Met His Thr Ser Leu Ile Leu
    2610   2615   2620
Glu Lys Glu Gln Val Glu Ile Ala Glu Lys Asn Val Leu Glu Lys Glu
2625   2630   2635   2640
Lys Lys Leu Leu Glu Leu Gln Lys Leu Leu Glu Gly Asn Glu Lys Lys
            2645   2650   2655
Gln Arg Glu Lys Glu Lys Lys Arg Ser Pro Gln Asp Val Glu Val Leu
        2660   2665   2670
Lys Thr Thr Thr Glu Leu Phe His Ser Asn Glu Glu Ser Gly Phe Phe

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                2675                    2680                    2685
Asn Glu Leu Glu Ala Leu Arg Ala Glu Ser Val Ala Thr Lys Ala Glu
    2690                    2695                    2700
Leu Ala Ser Tyr Lys Glu Lys Ala Glu Lys Leu Gln Glu Glu Leu Leu
2705                    2710                    2715                    2720
Val Lys Glu Thr Asn Met Thr Ser Leu Gln Lys Asp Leu Ser Gln Val
                2725                    2730                    2735
Arg Asp His Leu Ala Glu Ala Lys Glu Lys Leu Ser Ile Leu Glu Lys
            2740                    2745                    2750
Glu Asp Glu Thr Glu Val Gln Glu Ser Lys Lys Ala Cys Met Phe Glu
            2755                    2760                    2765
Pro Leu Pro Ile Lys Leu Ser Lys Ser Ile Ala Ser Gln Thr Asp Gly
        2770                    2775                    2780
Thr Leu Lys Ile Ser Ser Ser Asn Gln Thr Pro Gln Ile Leu Val Lys
2785                    2790                    2795                    2800
Asn Ala Gly Ile Gln Ile Asn Leu Gln Ser Glu Cys Ser Ser Glu Glu
                2805                    2810                    2815
Val Thr Glu Ile Ile Ser Gln Phe Thr Glu Lys Ile Glu Lys Met Gln
            2820                    2825                    2830
Glu Leu His Ala Ala Glu Ile Leu Asp Met Glu Ser Arg His Ile Ser
            2835                    2840                    2845
Glu Thr Glu Thr Leu Lys Arg Glu His Tyr Val Ala Val Gln Leu Leu
    2850                    2855                    2860
Lys Glu Glu Cys Gly Thr Leu Lys Ala Val Ile Gln Cys Leu Arg Ser
2865                    2870                    2875                    2880
Lys Glu Gly Ser Ser Ile Pro Glu Leu Ala His Ser Asp Ala Tyr Gln
                2885                    2890                    2895
Thr Arg Glu Ile Cys Ser Ser Asp Ser Gly Ser Asp Trp Gly Gln Gly
            2900                    2905                    2910
Ile Tyr Leu Thr His Ser Gln Gly Phe Asp Ile Ala Ser Glu Gly Arg
            2915                    2920                    2925
Gly Glu Ser Glu Ser Ala Thr Asp Ser Phe Pro Lys Lys Ile Lys
    2930                    2935                    2940
Gly Leu Leu Arg Ala Val His Asn Glu Gly Met Gln Val Leu Ser Leu
2945                    2950                    2955                    2960
Thr Glu Ser Pro Tyr Ser Asp Gly Glu Asp His Ser Ile Gln Gln Val
                2965                    2970                    2975
Ser Glu Pro Trp Leu Glu Glu Arg Lys Ala Tyr Ile Asn Thr Ile Ser
            2980                    2985                    2990
Ser Leu Lys Asp Leu Ile Thr Lys Met Gln Leu Gln Arg Glu Ala Glu
            2995                    3000                    3005
Val Tyr Asp Ser Ser Gln Ser His Glu Ser Phe Ser Asp Trp Arg Gly
        3010                    3015                    3020
Glu Leu Leu Leu Ala Leu Gln Gln Val Phe Leu Glu Glu Arg Ser Val
3025                    3030                    3035                    3040
Leu Leu Ala Ala Phe Arg Thr Glu Leu Thr Ala Leu Gly Thr Thr Asp
            3045                    3050                    3055
Ala Val Gly Leu Leu Asn Cys Leu Glu Gln Arg Ile Gln Glu Gln Gly
            3060                    3065                    3070
Val Glu Tyr Gln Ala Ala Met Glu Cys Leu Gln Lys Ala Asp Arg Arg
        3075                    3080                    3085
Ser Leu Leu Ser Glu Ile Gln Ala Leu His Ala Gln Met Asn Gly Arg
        3090                    3095                    3100
Lys Ile Thr Leu Lys Arg Glu Gln Glu Ser Glu Lys Pro Ser Gln Glu
3105                    3110                    3115                    3120
Leu Leu Glu Tyr Asn Ile Gln Gln Lys Gln Ser Gln Met Leu Glu Met
            3125                    3130                    3135
Gln Val Glu Leu Ser Ser Met Lys Asp Arg Ala Thr Glu Leu Gln Glu
        3140                    3145                    3150
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Gln Leu Ser Ser Glu Lys Met Val Val Ala Glu Leu Lys Ser Glu Leu
        3155            3160            3165
Ala Gln Thr Lys Leu Glu Leu Glu Thr Thr Leu Lys Ala Gln His Lys
        3170            3175            3180
His Leu Lys Glu Leu Glu Ala Phe Arg Leu Glu Val Lys Asp Lys Thr
3185            3190            3195            3200
Asp Glu Val His Leu Leu Asn Asp Thr Leu Ala Ser Glu Gln Lys Lys
            3205            3210            3215
Ser Arg Glu Leu Gln Trp Ala Leu Glu Lys Glu Lys Ala Lys Leu Gly
        3220            3225            3230
Arg Ser Glu Glu Arg Asp Lys Glu Glu Leu Glu Asp Leu Lys Phe Ser
        3235            3240            3245
Leu Glu Ser Gln Lys Gln Arg Asn Leu Gln Leu Asn Leu Leu Leu Glu
        3250            3255            3260
Gln Gln Lys Gln Leu Leu Asn Glu Ser Gln Gln Lys Ile Glu Ser Gln
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38

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gccaccttgg ccctgcttgc ccggatgggg gggcagccag ctttcacgga tctagaggtg 11460
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atttccagaa tgaaattttt ggttcgacgg tggcatcgag tcacaggttc tgtttccatc 11580
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cattcttctg gtgggctgga gttatatgga gaaccaagac atactacgta tcgctcaaga 11700
tcagatctgg actatattag gtcccctta ccatttcaga ataggtaccc aggcactcca 11760
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gccctaacag attatatcac tcggctagag gcactgcaaa gacgacttgg aactatacag 11880
tcaggtgctc tgagtttaac cacatcttgg cagcaccaca gtgcgagacc cacagctccc 11940
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aataatttaa gtgaaaatat gatttatcac cccagatccc antcctccca aaaatgattt 12120
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ggtactctga gtttacagag cctaactgga gaacgtattc ctaagtagcg catggcagaa 12240
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gaaaaaaaaa aaa                                                      12313
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<210> 8
<211> 3917
<212> PRT
<213> Homo sapiens

<400> 8

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Met Glu Asp Glu Glu Arg Gln Lys Lys Leu Glu Ala Gly Lys Ala Lys
1               5                   10                  15
Leu Ala Gln Phe Arg Gln Arg Lys Ala Gln Ser Asp Gly Gln Ser Pro
            20                  25                  30
Ser Lys Lys Gln Lys Lys Lys Arg Lys Thr Ser Ser Ser Lys His Asp
        35                  40                  45
Val Ser Ala His His Asp Leu Asn Ile Asp Gln Ser Gln Cys Asn Glu
    50                  55                  60
Met Tyr Ile Asn Ser Ser Gln Arg Val Glu Ser Thr Val Ile Pro Glu
65                  70                  75                  80
Ser Thr Ile Met Arg Thr Leu His Ser Gly Glu Ile Thr Ser His Glu
                85                  90                  95
Gln Gly Phe Ser Val Glu Leu Glu Ser Glu Ile Ser Thr Thr Ala Asp
            100                 105                 110
Asp Cys Ser Ser Glu Val Asn Gly Cys Ser Phe Val Met Arg Thr Gly
        115                 120                 125
Lys Pro Thr Asn Leu Leu Arg Glu Glu Glu Phe Gly Val Asp Asp Ser
    130                 135                 140
Tyr Ser Glu Gln Gly Ala Gln Asp Ser Pro Thr His Leu Glu Met Met
145                 150                 155                 160
Glu Ser Glu Leu Ala Gly Lys Gln His Glu Ile Glu Glu Leu Asn Arg
                165                 170                 175
Glu Leu Glu Glu Met Arg Val Thr Tyr Gly Thr Glu Gly Leu Gln Gln
            180                 185                 190
Leu Gln Glu Phe Glu Ala Ala Ile Lys Gln Arg Asp Gly Ile Ile Thr
        195                 200                 205
Gln Leu Thr Ala Asn Leu Gln Gln Ala Arg Arg Glu Lys Asp Glu Thr
    210                 215                 220
Met Arg Glu Phe Leu Glu Leu Thr Glu Gln Ser Gln Lys Leu Gln Ile
225                 230                 235                 240
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Gln Phe Gln Gln Leu Gln Ala Ser Glu Thr Leu Arg Asn Ser Thr His
            245             250             255
Ser Ser Thr Ala Ala Asp Leu Leu Gln Ala Lys Gln Gln Ile Leu Thr
            260             265             270
His Gln Gln Gln Leu Glu Glu Gln Asp His Leu Leu Glu Asp Tyr Gln
            275             280             285
Lys Lys Lys Glu Asp Phe Thr Met Gln Ile Ser Phe Leu Gln Glu Lys
        290             295             300
Ile Lys Val Tyr Glu Met Glu Gln Asp Lys Lys Val Glu Asn Ser Asn
305             310             315             320
Lys Glu Glu Ile Gln Glu Lys Glu Thr Ile Ile Glu Glu Leu Asn Thr
            325             330             335
Lys Ile Ile Glu Glu Glu Lys Lys Thr Leu Glu Leu Lys Asp Lys Leu
            340             345             350
Thr Thr Ala Asp Lys Leu Leu Gly Glu Leu Gln Glu Gln Ile Val Gln
            355             360             365
Lys Asn Gln Glu Ile Lys Asn Met Lys Leu Glu Leu Thr Asn Ser Lys
        370             375             380
Gln Lys Glu Arg Gln Ser Ser Glu Glu Ile Lys Gln Leu Met Gly Thr
385             390             395             400
Val Glu Glu Leu Gln Lys Arg Asn His Lys Asp Ser Gln Phe Glu Thr
            405             410             415
Asp Ile Val Gln Arg Met Glu Gln Glu Thr Gln Arg Lys Leu Glu Gln
            420             425             430
Leu Arg Ala Glu Leu Asp Glu Met Tyr Gly Gln Gln Ile Val Gln Met
            435             440             445
Lys Gln Glu Leu Ile Arg Gln His Met Ala Gln Met Glu Glu Met Lys
        450             455             460
Thr Arg His Lys Gly Glu Met Glu Asn Ala Leu Arg Ser Tyr Ser Asn
465             470             475             480
Ile Thr Val Asn Glu Asp Gln Ile Lys Leu Met Asn Val Ala Ile Asn
            485             490             495
Glu Leu Asn Ile Lys Leu Gln Asp Thr Asn Ser Gln Lys Glu Lys Leu
            500             505             510
Lys Glu Glu Leu Gly Leu Ile Leu Glu Glu Lys Cys Ala Leu Gln Arg
            515             520             525
Gln Leu Glu Asp Leu Val Glu Glu Leu Ser Phe Ser Arg Glu Gln Ile
            530             535             540
Gln Arg Ala Arg Gln Thr Ile Ala Glu Gln Glu Ser Lys Leu Asn Glu
545             550             555             560
Ala His Lys Ser Leu Ser Thr Val Glu Asp Leu Lys Ala Glu Ile Val
            565             570             575
Ser Ala Ser Glu Ser Arg Lys Glu Leu Glu Leu Lys His Glu Ala Glu
            580             585             590
Val Thr Asn Tyr Lys Ile Lys Leu Glu Met Leu Glu Lys Glu Lys Asn
            595             600             605
Ala Val Leu Asp Arg Met Ala Glu Ser Gln Glu Ala Glu Leu Glu Arg
            610             615             620
Leu Arg Thr Gln Leu Leu Phe Ser His Glu Glu Glu Leu Ser Lys Leu
625             630             635             640
Lys Glu Asp Leu Glu Ile Glu His Arg Ile Asn Ile Glu Lys Leu Lys
            645             650             655
Asp Asn Leu Gly Ile His Tyr Lys Gln Ile Asp Gly Leu Gln Asn
            660             665             670
Glu Met Ser Gln Lys Ile Glu Thr Met Gln Phe Glu Lys Asp Asn Leu
            675             680             685
Ile Thr Lys Gln Asn Gln Leu Ile Leu Glu Ile Ser Lys Leu Lys Asp
        690             695             700
Leu Gln Gln Ser Leu Val Asn Ser Lys Ser Glu Glu Met Thr Leu Gln
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705                     710                     715                     720
Ile Asn Glu Leu Gln Lys Glu Ile Glu Ile Leu Arg Gln Glu Glu Lys
                    725          ·          730              735
Glu Lys Gly Thr Leu Glu Gln Glu Val Gln Glu Leu Gln Leu Lys Thr
            740                     745                 750
Glu Leu Leu Glu Lys Gln Met Lys Glu Lys Glu Asn Asp Leu Gln Glu
        755                     760                     765
Lys Phe Ala Gln Leu Glu Ala Glu Asn Ser Ile Leu Lys Asp Glu Lys
    770                     775                     780
Lys Thr Leu Glu Asp Met Leu Lys Ile His Thr Pro Val Ser Gln Glu
·785                     790                     795                 800
Glu Arg Leu Ile Phe Leu Asp Ser Ile Lys Ser Lys Ser Lys Asp Ser
                805                     810                     815
Val Trp Glu Lys Glu Ile Glu Ile Leu Ile Glu Glu Asn Glu Asp Leu
    ·           820                     825                 830
Lys Gln Gln Cys Ile Gln Leu Asn Glu Glu Ile Glu Lys Gln Arg Asn
            835                     840                     845
Thr Phe Ser Phe Ala Glu Lys Asn Phe Glu Val Asn Tyr Gln Glu Leu
    850                     855                     860
Gln Glu Glu Tyr Ala Cys Leu Leu Lys Val Lys Asp Asp Leu Glu Asp
865                     870                     875                 880
Ser Lys Asn Lys Gln Glu Leu Glu Tyr Lys Ser Lys Leu Lys Ala Leu
        ·       885                     890                     895
Asn Glu Glu Leu His Leu Gln Arg Ile Asn Pro Thr Thr Val Lys Met
            900                     905                     910
Lys Ser Ser Val Phe Asp Glu Asp Lys Thr Phe Val Ala Glu Thr Leu
            915                     920                 925
Glu Met Gly Glu Val Val Glu Lys Asp Thr Thr Glu Leu Met Glu Lys
    930                     935                     940
Leu Glu Val Thr Lys Arg Glu Lys Leu Glu Leu Ser Gln Arg Leu Ser
945                     950                     955                 960
Asp Leu Ser Glu Gln Leu Lys Gln Lys His Gly Glu Ile Ser Phe Leu
                965                     970                     975
Asn Glu ·Glu Val Lys Ser Leu Lys Gln Glu Lys Glu Gln Val Ser Leu
            980                     985                     990
Arg Cys Arg Glu Leu Glu Ile Ile Ile Asn His Asn Arg Ala Glu Asn
        995                     1000                    1005
Val Gln Ser Cys Asp Thr Gln Val Ser Ser Leu Leu Asp Gly Val Val
    1010                    1015                    1020
Thr Met Thr Ser Arg Gly Ala Glu Gly Ser Val Ser Lys Val Asn Lys
1025                    1030                    1035                1040
Ser Phe Gly Glu Glu Ser Lys Ile Met Val Glu Asp Lys Val Ser Phe
                1045                    1050                    1055
Glu Asn Met Thr Val Gly Glu Glu Ser Lys Gln·Glu Gln Leu Ile Leu
            1060                    1065                    1070
Asp His Leu Pro Ser Val Thr Lys Glu Ser Ser Leu Arg Ala Thr Gln
    ·       1075                    1080                    1085
Pro Ser Glu Asn Asp Lys Leu Gln Lys Glu Leu Asn Val Leu Lys Ser
    1090                    1095                    1100
Glu Gln Asn Asp Leu Arg Leu Gln Met Glu Ala Gln Arg Ile Cys Leu
1105                    1110                    1115                1120
Ser Leu Val Tyr Ser Thr His Val Asp Gln Val Arg Glu Tyr Met Glu
                1125                    1130                    1135
Asn Glu Lys Asp Lys Ala Leu Cys Ser Leu Lys Glu Glu Leu Ile Phe
            1140                    1145                    1150
Ala Gln Glu Glu Lys Ile Lys Glu Leu Gln Lys Ile His Gln Leu Glu
        1155                    1160                    1165
Leu Gln Thr Met Lys Thr Gln Glu Thr Gly Asp Glu Gly Lys Pro Leu
            1170                    1175                    1180
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His Leu Leu Ile Gly Lys Leu Gln Lys Ala Val Ser Glu Glu Cys Ser
1185                1190                1195                1200
Tyr Phe Leu Gln Thr Leu Cys Ser Val Leu Gly Glu Tyr Tyr Thr Pro
               1205                1210                1215
Ala Leu Lys Cys Glu Val Asn Ala Glu Asp Lys Glu Asn Ser Gly Asp
               1220                1225                1230
Tyr Ile Ser Glu Asn Glu Asp Pro Glu Leu Gln Asp Tyr Arg Tyr Glu
               1235                1240                1245
Val Gln Asp Phe Gln Glu Asn Met His Thr Leu Leu Asn Lys Val Thr
        1250                1255                1260
Glu Glu Tyr Asn Lys Leu Leu Val Leu Gln Thr Arg Leu Ser Lys Ile
1265                1270                1275                1280
Trp Gly Gln Gln Thr Asp Gly Met Lys Leu Glu Phe Gly Glu Glu Asn
               1285                1290                1295
Leu Pro Lys Glu Glu Thr Glu Phe Leu Ser Ile His Ser Gln Met Thr
               1300                1305                1310
Asn Leu Glu Asp Ile Asp Val Asn His Lys Ser Lys Leu Ser Ser Leu
               1315                1320                1325
Gln Asp Leu Glu Lys Thr Lys Leu Glu Glu Gln Val Gln Glu Leu Glu
        1330                1335                1340
Ser Leu Ile Ser Ser Leu Gln Gln Gln Leu Lys Glu Thr Glu Gln Asn
1345                1350                1355                1360
Tyr Glu Ala Glu Ile His Cys Leu Gln Lys Arg Leu Gln Ala Val Ser
               1365                1370                1375
Glu Ser Thr Val Pro Pro Ser Leu Pro Val Asp Ser Val Val Ile Thr
               1380                1385                1390
Glu Ser Asp Ala Gln Arg Thr Met Tyr Pro Gly Ser Cys Val Lys Lys
        1395                1400                1405
Asn Ile Asp Gly Thr Ile Glu Phe Ser Gly Glu Phe Gly Val Lys Glu
        1410                1415                1420
Glu Thr Asn Ile Val Lys Leu Leu Glu Lys Gln Tyr Gln Glu Gln Leu
1425                1430                1435                1440
Glu Glu Glu Val Ala Lys Val Ile Val Ser Met Ser Ile Ala Phe Ala
               1445                1450                1455
Gln Gln Thr Glu Leu Ser Arg Ile Ser Gly Gly Lys Glu Asn Thr Ala
        1460                1465                1470
Ser Ser Lys Gln Ala His Ala Val Cys Gln Gln Glu Gln His Tyr Phe
        1475                1480                1485
Asn Glu Met Lys Leu Ser Gln Asp Gln Ile Gly Phe Gln Thr Phe Glu
        1490                1495                1500
Thr Val Asp Val Lys Phe Lys Glu Glu Phe Lys Pro Leu Ser Lys Glu
1505                1510                1515                1520
Leu Gly Glu His Gly Lys Glu Ile Leu Leu Ser Asn Ser Asp Pro His
               1525                1530                1535
Asp Ile Pro Glu Ser Lys Asp Cys Val Leu Thr Ile Ser Glu Glu Met
        1540                1545                1550
Phe Ser Lys Asp Lys Thr Phe Ile Val Arg Gln Ser Ile His Asp Glu
        1555                1560                1565
Ile Ser Val Ser Ser Met Asp Ala Ser Arg Gln Leu Met Leu Asn Glu
        1570                1575                1580
Glu Gln Leu Glu Asp Met Arg Gln Glu Leu Val Arg Gln Tyr Gln Glu
1585                1590                1595                1600
His Gln Gln Ala Thr Glu Leu Leu Arg Gln Ala His Met Arg Gln Met
               1605                1610                1615
Glu Arg Gln Arg Glu Asp Gln Glu Gln Leu Gln Glu Glu Ile Lys Arg
        1620                1625                1630
Leu Asn Arg Gln Leu Ala Gln Arg Ser Ser Ile Asp Asn Glu Asn Leu
        1635                1640                1645
Val Ser Glu Arg Glu Arg Val Leu Leu Glu Glu Leu Glu Ala Leu Lys
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        1650              1655                  1660
Gln Leu Ser Leu Ala Gly Arg Glu Lys Leu Cys Cys Glu Leu Arg Asn
1665              1670                  1675              1680
Ser Ser Thr Gln Thr Gln Asn Gly Asn Glu Asn Gln Gly Glu Val Glu
                  1685              1690              1695
Glu Gln Thr Phe Lys Glu Lys Glu Leu Asp Arg Lys Pro Glu Asp Val
              1700                  1705              1710
Pro Pro Glu Ile Leu Ser Asn Glu Arg Tyr Ala Leu Gln Lys Ala Asn
              1715                  1720              1725
Asn Arg Leu Leu Lys Ile Leu Leu Glu Val Val Lys Thr Thr Ala Ala
          1730                  1735              1740
Val Glu Glu Thr Ile Gly Arg His Val Leu Gly Ile Leu Asp Arg Ser
1745                  ·1750                  1755              1760
Ser Lys Ser Gln Ser Ser Ala Ser Leu Ile Trp Arg Ser Glu Ala Glu.
              1765                  1770              1775
Ala Ser Val Lys Ser Cys Val His Glu Glu His Thr Arg Val Thr Asp
          1780                  1785              1790
Glu Ser Ile Pro Ser Tyr Ser Gly Ser Asp Met Pro Arg Asn Asp Ile
          1795                  1800              1805
Asn Met Trp Ser Lys Val Thr Glu Glu Gly Thr Glu Leu Ser Gln Arg
      1810                  1815              1820
Leu Val Arg Ser Gly Phe Ala Gly Thr Glu Ile Asp Pro Glu Asn Glu
1825                  1830                  1835              1840
Glu Leu Met Leu Asn Ile Ser Ser Arg Leu Gln Ala Ala Val Glu Lys
              1845                  1850              1855
Leu Leu Glu Ala Ile Ser Glu Thr Ser Ser Gln Leu Glu His Ala Lys
      1860                  1865              1870
Val Thr Gln Thr Glu Leu Met Arg Glu Ser Phe Arg Gln Lys Gln Glu
          1875                  1880              1885
Ala Thr Glu Ser Leu Lys Cys Gln Glu Glu Leu Arg Glu Arg Leu His
      1890                  1895              1900
Glu Glu Ser Arg Ala Arg Glu Gln Leu Ala Val Glu Leu Ser Lys Ala
1905                  1910                  1915              1920
Glu Gly Val Ile Asp Gly Tyr Ala Asp Glu Lys Thr Leu Phe Glu Arg
              1925                  1930              1935
Gln Ile Gln Glu Lys Thr Asp Ile Ile Asp Arg Leu Glu Gln Glu Leu
          1940                  1945              1950
Leu Cys Ala Ser Asn Arg Leu Gln Glu Leu Glu Ala Glu Gln Gln Gln
          1955                  1960              1965
Ile Gln Glu Glu Arg Glu Leu Leu Ser Arg Gln Lys Glu Ala Met Lys
      1970                  1975              1980
Ala Glu Ala Gly Pro Val Glu Gln Gln Leu Leu Gln Glu Thr Glu Lys
1985                  1990                  1995              2000
Leu Met Lys Glu Lys Leu Glu Val Gln Cys Gln Ala Glu Lys Val Arg
              2005                  2010              2015
Asp Asp Leu Gln Lys Gln Val Lys Ala Leu Glu Ile Asp Val Glu Glu
          2020                  2025              2030
Gln Val Ser Arg Phe Ile Glu Leu Glu Gln Glu Lys Asn Thr Glu Leu
          2035                  2040              2045
Met Asp Leu Arg Gln Gln Asn Gln Ala Leu Glu Lys Gln Leu Glu Lys
      2050                  2055              2060
Met Arg Lys Phe Leu Asp Glu Gln Ala Ile Asp Arg Glu His Glu Arg
2065                  2070                  2075              2080
Asp Val Phe Gln Gln Glu Ile Gln Lys Leu Glu Gln Gln Leu Lys Val
              2085                  2090              2095
Val Pro Arg Phe Gln Pro Ile Ser Glu His Gln Thr Arg Glu Val Glu
          2100                  2105              2110
Gln Leu Ala Asn His Leu Lys Glu Lys Thr Asp Lys Cys Ser Glu Leu
      2115                  2120              2125
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Leu Leu Ser Lys Glu Gln Leu Gln Arg Asp Ile Gln Glu Arg Asn Glu
    2130            2135            2140
Glu Ile Glu Lys Leu Glu Phe Arg Val Arg Glu Leu Glu Gln Ala Leu
2145            2150            2155            2160
Leu Val Glu Asp Arg Lys His Phe Gly Ala Val Glu Ala Lys Pro Glu
            2165            2170            2175
Leu Ser Leu Glu Val Gln Leu Gln Ala Glu Arg Asp Ala Ile Asp Arg
            2180            2185            2190
Lys Glu Lys Glu Ile Thr Asn Leu Glu Glu Gln Leu Glu Gln Phe Arg
        2195            2200            2205
Glu Glu Leu Glu Asn Lys Asn Glu Glu Val Gln Gln Leu His Met Gln
        2210            2215            2220
Leu Glu Ile Gln Lys Lys Glu Ser Thr Thr Arg Leu Gln Glu Leu Glu
2225            2230            2235            2240
Gln Glu Asn Lys Leu Phe Lys Asp Asp Met Glu Lys Leu Gly Leu Ala
            2245            2250            2255
Ile Lys Glu Ser Asp Ala Met Ser Thr Gln Asp Gln His Val Leu Phe
        2260            2265            2270
Gly Lys Phe Ala Gln Ile Ile Gln Glu Lys Glu Val Glu Ile Asp Gln
        2275            2280            2285
Leu Asn Glu Gln Val Thr Lys Leu Gln Gln Gln Leu Lys Ile Thr Thr
    2290            2295            2300
Asp Asn Lys Val Ile Glu Glu Lys Asn Glu Leu Ile Arg Asp Leu Glu
2305            2310            2315            2320
Thr Gln Ile Glu Cys Leu Met Ser Asp Gln Glu Cys Val Lys Arg Asn
            2325            2330            2335
Arg Glu Glu Glu Ile Glu Gln Leu Asn Glu Val Ile Glu Lys Leu Gln
            2340            2345            2350
Gln Glu Leu Ala Asn Ile Gly Gln Lys Thr Ser Met Asn Ala His Ser
        2355            2360            2365
Leu Ser Glu Glu Ala Asp Ser Leu Lys His Gln Leu Asp Val Val Ile
    2370            2375            2380
Ala Glu Lys Leu Ala Leu Glu Gln Gln Val Glu Thr Ala Asn Glu Glu
2385            2390            2395            2400
Met Thr Phe Met Lys Asn Val Leu Lys Glu Thr Asn Phe Lys Met Asn
            2405            2410            2415
Gln Leu Thr Gln Glu Leu Phe Ser Leu Lys Arg Glu Arg Glu Ser Val
        2420            2425            2430
Glu Lys Ile Gln Ser Ile Pro Glu Asn Ser Val Asn Val Ala Ile Asp
        2435            2440            2445
His Leu Ser Lys Asp Lys Pro Glu Leu Glu Val Val Leu Thr Glu Asp
    2450            2455            2460
Ala Leu Lys Ser Leu Glu Asn Gln Thr Tyr Phe Lys Ser Phe Glu Glu
2465            2470            2475            2480
Asn Gly Lys Gly Ser Ile Ile Asn Leu Glu Thr Arg Leu Leu Gln Leu
            2485            2490            2495
Glu Ser Thr Val Ser Ala Lys Asp Leu Glu Leu Thr Gln Cys Tyr Lys
        2500            2505            2510
Gln Ile Lys Asp Met Gln Glu Gln Gly Gln Phe Glu Thr Glu Met Leu
    2515            2520            2525
Gln Lys Lys Ile Val Asn Leu Gln Lys Ile Val Glu Glu Lys Val Ala
    2530            2535            2540
Ala Ala Leu Val Ser Gln Ile Gln Leu Glu Ala Val Gln Glu Tyr Ala
2545            2550            2555            2560
Lys Phe Cys Gln Asp Asn Gln Thr Ile Ser Ser Glu Pro Glu Arg Thr
            2565            2570            2575
Asn Ile Gln Asn Leu Asn Gln Leu Arg Glu Asp Glu Leu Gly Ser Asp
            2580            2585            2590
Ile Ser Ala Leu Thr Leu Arg Ile Ser Glu Leu Glu Ser Gln Val Val
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              2595                2600                2605
Glu Met His Thr Ser Leu Ile Leu Glu Lys Glu Gln Val Glu Ile Ala
      2610                2615                2620
Glu Lys Asn Val Leu Glu Lys Glu Lys Lys Leu Leu Glu Leu Gln Lys
2625                2630                2635                2640
Leu Leu Glu Gly Asn Glu Lys Lys Gln Arg Glu Lys Glu Lys Lys Arg
              2645                2650                2655
Ser Pro Gln Asp Val Glu Val Leu Lys Thr Thr Thr Glu Leu Phe His
              2660                2665                2670
Ser Asn Glu Glu Ser Gly Phe Phe Asn Glu Leu Glu Ala Leu Arg Ala
              2675                2680                2685
Glu Ser Val Ala Thr Lys Ala Glu Leu Ala Ser Tyr Lys Glu Lys Ala
              2690                2695                2700
Glu Lys Leu Gln Glu Glu Leu Leu Val Lys Glu Thr Asn Met Thr Ser
2705                2710                2715                2720
Leu Gln Lys Asp Leu Ser Gln Val Arg Asp His Leu Ala Glu Ala Lys
              2725                2730                2735
Glu Lys Leu Ser Ile Leu Glu Lys Glu Asp Glu Thr Glu Val Gln Glu
              2740                2745                2750
Ser Lys Lys Ala Cys Met Phe Glu Pro Leu Pro Ile Lys Leu Ser Lys
              2755                2760                2765
Ser Ile Ala Ser Gln Thr Asp Gly Thr Leu Lys Ile Ser Ser Ser Asn
      2770                2775                2780
Gln Thr Pro Gln Ile Leu Val Lys Asn Ala Gly Ile Gln Ile Asn Leu
2785                2790                2795                2800
Gln Ser Glu Cys Ser Ser Glu Glu Val Thr Glu Ile Ile Ser Gln Phe
              2805                2810                2815
Thr Glu Lys Ile Glu Lys Met Gln Glu Leu His Ala Ala Glu Ile Leu
      2820                2825                2830
Asp Met Glu Ser Arg His Ile Ser Glu Thr Glu Thr Leu Lys Arg Glu
      2835                2840                2845
His Tyr Val Ala Val Gln Leu Leu Lys Glu Glu Cys Gly Thr Leu Lys
      2850                2855                2860
Ala Val Ile Gln Cys Leu Arg Ser Lys Glu Gly Ser Ser Ile Pro Glu
2865                2870                2875                2880
Leu Ala His Ser Asp Ala Tyr Gln Thr Arg Glu Ile Cys Ser Ser Asp
              2885                2890                2895
Ser Gly Ser Asp Trp Gly Gln Gly Ile Tyr Leu Thr His Ser Gln Gly
      2900                2905                2910
Phe Asp Ile Ala Ser Glu Gly Arg Gly Glu Glu Ser Glu Ser Ala Thr
      2915                2920                2925
Asp Ser Phe Pro Lys Lys Ile Lys Gly Leu Leu Arg Ala Val His Asn
      2930                2935                2940
Glu Gly Met Gln Val Leu Ser Leu Thr Glu Ser Pro Tyr Ser Asp Gly
2945                2950                2955                2960
Glu Asp His Ser Ile Gln Gln Val Ser Glu Pro Trp Leu Glu Glu Arg
              2965                2970                2975
Lys Ala Tyr Ile Asn Thr Ile Ser Ser Leu Lys Asp Leu Ile Thr Lys
              2980                2985                2990
Met Gln Leu Gln Arg Glu Ala Glu Val Tyr Asp Ser Ser Gln Ser His
              2995                3000                3005
Glu Ser Phe Ser Asp Trp Arg Gly Glu Leu Leu Leu Ala Leu Gln Gln
      3010                3015                3020
Val Phe Leu Glu Glu Arg Ser Val Leu Leu Ala Ala Phe Arg Thr Glu
3025                3030                3035                3040
Leu Thr Ala Leu Gly Thr Thr Asp Ala Val Gly Leu Leu Asn Cys Leu
              3045                3050                3055
Glu Gln Arg Ile Gln Glu Gln Gly Val Glu Tyr Gln Ala Ala Met Glu
              3060                3065                3070
```

Cys Leu Gln Lys Ala Asp Arg Arg Ser Leu Leu Ser Glu Ile Gln Ala
        3075   3080   3085
Leu His Ala Gln Met Asn Gly Arg Lys Ile Thr Leu Lys Arg Glu Gln
        3090   3095   3100
Glu Ser Glu Lys Pro Ser Gln Glu Leu Leu Glu Tyr Asn Ile Gln Gln
3105   3110   3115   3120
Lys Gln Ser Gln Met Leu Glu Met Gln Val Glu Leu Ser Ser Met Lys
                3125   3130   3135
Asp Arg Ala Thr Glu Leu Gln Glu Gln Leu Ser Ser Glu Lys Met Val
                3140   3145   3150
Val Ala Glu Leu Lys Ser Glu Leu Ala Gln Thr Lys Leu Glu Leu Glu
        3155   3160   3165
Thr Thr Leu Lys Ala Gln His Lys His Leu Lys Glu Leu Glu Ala Phe
        3170   3175   3180
Arg Leu Glu Val Lys Asp Lys Thr Asp Glu Val His Leu Leu Asn Asp
3185   3190   3195   3200
Thr Leu Ala Ser Glu Gln Lys Lys Ser Arg Glu Leu Gln Trp Ala Leu
                3205   3210   3215
Glu Lys Glu Lys Ala Lys Leu Gly Arg Ser Glu Glu Arg Asp Lys Glu
                3220   3225   3230
Glu Leu Glu Asp Leu Lys Phe Ser Leu Glu Ser Gln Lys Gln Arg Asn
        3235   3240   3245
Leu Gln Leu Asn Leu Leu Leu Glu Gln Gln Lys Gln Leu Leu Asn Glu
        3250   3255   3260
Ser Gln Gln Lys Ile Glu Ser Gln Arg Met Leu Tyr Asp Ala Gln Leu
3265   3270   3275   3280
Ser Glu Glu Gln Gly Arg Asn Leu Glu Leu Gln Val Leu Leu Glu Ser
                3285   3290   3295
Glu Lys Val Arg Ile Arg Glu Met Ser Ser Thr Leu Asp Arg Glu Arg
        3300   3305   3310
Glu Leu His Ala Gln Leu Gln Ser Ser Asp Gly Thr Gly Gln Ser Arg
        3315   3320   3325
Pro Pro Leu Pro Ser Glu Asp Leu Leu Lys Glu Leu Gln Lys Gln Leu
        3330   3335   3340
Glu Glu Lys His Ser Arg Ile Val Glu Leu Leu Asn Glu Thr Glu Lys
3345   3350   3355   3360
Tyr Lys Leu Asp Ser Leu Gln Thr Arg Gln Gln Met Glu Lys Asp Arg
                3365   3370   3375
Gln Val His Arg Lys Thr Leu Gln Thr Glu Gln Glu Ala Asn Thr Glu
        3380   3385   3390
Gly Gln Lys Lys Met His Glu Leu Gln Ser Lys Val Glu Asp Leu Gln
        3395   3400   3405
Arg Gln Leu Glu Glu Lys Arg Gln Gln Val Tyr Lys Leu Asp Leu Glu
        3410   3415   3420
Gly Gln Arg Leu Gln Gly Ile Met Gln Glu Phe Gln Lys Gln Glu Leu
3425   3430   3435   3440
Glu Arg Glu Glu Lys Arg Glu Ser Arg Arg Ile Leu Tyr Gln Asn Leu
                3445   3450   3455
Asn Glu Pro Thr Thr Trp Ser Leu Thr Ser Asp Arg Thr Arg Asn Trp
        3460   3465   3470
Val Leu Gln Gln Lys Ile Glu Gly Glu Thr Lys Glu Ser Asn Tyr Ala
        3475   3480   3485
Lys Leu Ile Glu Met Asn Gly Gly Gly Thr Gly Cys Asn His Glu Leu
        3490   3495   3500
Glu Met Ile Arg Gln Lys Leu Gln Cys Val Ala Ser Lys Leu Gln Val
3505   3510   3515   3520
Leu Pro Gln Lys Ala Ser Glu Arg Leu Gln Phe Glu Thr Ala Asp Asp
                3525   3530   3535
Glu Asp Phe Ile Trp Val Gln Glu Asn Ile Asp Glu Ile Ile Leu Gln

```
                    3540              3545                3550
     Leu Gln Lys Leu Thr Gly Gln Gln Gly Glu Glu Pro Ser Leu Val Ser
           3555              3560              3565
     Pro Ser Thr Ser Cys Gly Ser Leu Thr Glu Arg Leu Leu Arg Gln Asn
           3570              3575              3580
     Ala Glu Leu Thr Gly His Ile Ser Gln Leu Thr Glu Glu Lys Asn Asp
     3585              3590              3595              3600
     Leu Arg Asn Met Val Met Lys Leu Glu Glu Gln Ile Arg Trp Tyr Arg
                    3605              3610              3615
     Gln Thr Gly Ala Gly Arg Asp Asn Ser Ser Arg Phe Ser Leu Asn Gly
                    3620              3625              3630
     Gly Ala Asn Ile Glu Ala Ile Ile Ala Ser Glu Lys Glu Val Trp Asn
                    3635              3640              3645
     Arg Glu Lys Leu Thr Leu Gln Lys Ser Leu Lys Arg Ala Glu Ala Glu
                    3650              3655              3660
     Val Tyr Lys Leu Lys Ala Glu Leu Arg Asn Asp Ser Leu Leu Gln Thr
     3665              3670              3675              3680
     Leu Ser Pro Asp Ser Glu His Val Thr Leu Lys Arg Ile Tyr Gly Lys
                    3685              3690              3695
     Tyr Leu Arg Ala Glu Ser Phe Arg Lys Ala Leu Ile Tyr Gln Lys Lys
                    3700              3705              3710
     Tyr Leu Leu Leu Leu Leu Gly Gly Phe Gln Glu Cys Glu Asp Ala Thr
                    3715              3720              3725
     Leu Ala Leu Leu Ala Arg Met Gly Gly Gln Pro Ala Phe Thr Asp Leu
           3730              3735              3740
     Glu Val Ile Thr Asn Arg Pro Lys Gly Phe Thr Arg Phe Arg Ser Ala
     3745              3750              3755              3760
     Val Arg Val Ser Ile Ala Ile Ser Arg Met Lys Phe Leu Val Arg Arg
                    3765              3770              3775
     Trp His Arg Val Thr Gly Ser Val Ser Ile Asn Ile Asn Arg Asp Gly
                    3780              3785              3790
     Phe Gly Leu Asn Gln Gly Ala Glu Lys Thr Asp Ser Phe Tyr His Ser
                    3795              3800              3805
     Ser Gly Gly Leu Glu Leu Tyr Gly Glu Pro Arg His Thr Thr Tyr Arg
           3810              3815              3820
     Ser Arg Ser Asp Leu Asp Tyr Ile Arg Ser Pro Leu Pro Phe Gln Asn
     3825              3830              3835              3840
     Arg Tyr Pro Gly Thr Pro Ala Asp Phe Asn Pro Gly Ser Leu Ala Cys
                    3845              3850              3855
     Ser Gln Leu Gln Asn Tyr Asp Pro Asp Arg Ala Leu Thr Asp Tyr Ile
                    3860              3865              3870
     Thr Arg Leu Glu Ala Leu Gln Arg Arg Leu Gly Thr Ile Gln Ser Gly
                    3875              3880              3885
     Ala Leu Ser Leu Thr Thr Ser Trp Gln His His Ser Ala Arg Pro Thr
     3890              3895              3900
     Ala Pro Leu Phe Phe Glu Ile Leu Ser His Ser Leu Gly
     3905              3910              3915
```

<210> 9
<211> 2850
<212> DNA
<213> Homo sapiens

<400> 9
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gttgtgactt tcccttcga  attcctcggt atatcttggg gactggagga cctgtctggt 60
tattatacag acgcataact ggaggtggga tccacacagc tcagaacagc tggatcttgc 120
tcagtctctg ccaggggaag attccttgga ggaggccctg cagcgacatg gagggagctg 180
ctttgctgag agtctctgtc ctctgcatct ggatgagtgc actttttcctt ggtgtgagag 240
```

```
tgagggcaga ggaagctgga gcgagggtgc aacaaaacgt tccaagtggg acagatactg 300
gagatcctca aagtaagccc ctcggtgact gggctgctgg caccatggac ccagagagca 360
gtatctttat tgaggatgcc attaagtatt tcaaggaaaa agtgagcaca cagaatctgc 420
tactcctgct gactgataat gaggcctgga acggattcgt ggctgctgct gaactgccca 480
ggaatgaggc agatgagctc cgtaaagctc tggacaacct tgcaagacaa atgatcatga 540
aagacaaaaa ctggcacgat aaaggccagc agtacagaaa ctggtttctg aaagagtttc 600
ctcggttgaa aagtaagctt gaggataaca taagaaggct ccgtgccctt gcagatgggg 660
ttcagaaggt ccacaaaggc accaccatcg ccaatgtggt gtctggctct ctcagcattt 720
cctctggcat cctgaccctc gtcggcatgg gtctggcacc cttcacagag ggaggcagcc 780
ttgtactctt ggaacctggg atggagttgg gaatcacagc cgctttgacc gggattacca 840
gcagtaccat agactacgga aagaagtggt ggacacaagc ccaagcccac gacctggtca 900
tcaaaagcct tgacaaattg aaggaggtga aggagttttt gggtgagaac atatccaact 960
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gcatcctgga aatgagcaga ggagtcaagc tcacggatgt ggcccctgta agcttctttc 1200
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tcaacaataa ttataagatt ctgcaggcgg accaagaact gtgaccacag ggcagggcag 1380
ccaccaggag agatatgcct ggcaggggcc aggacaaaat gcaaactttt ttttttttct 1440
gagacagagt cttgctctgt cgccaagttg gagtgcaatg gtgcgatctc agctcactgc 1500
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aagtgctggg attacaggcg tgagccatcg ctttttgaccc aaatgcaaac attttattag 1740
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atagctgagc acagcagggg aggggttaat gcagatggca gtgcagcaag gagaaggcag 1860
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agcagtttac tttagactaa agaatatatt gggggccggg gtgtagtggc tcatgcctgt 1980
aatccgagca ctttgggagg ccaaggcggg cggatcacga ggtcaggaga tcaagaccat 2040
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cgagactcca tctcaaaaaa aaaaaaaaaa agaatatatt gacggaagaa tagaccggag 2280
gcttgaagga accagcaatg agaaggccag gaaaagaaag agctgaaaat ggagaaagcc 2340
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caggttcaat gtcctccgaa gaatgaagtc tttccctggt gatggtcccc tgccctgtct 2460
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aacagtccca tcgctcttac ccggtaagta aacagtcaga aaattagcat gaaagcagtt 2760
tagcattggg aggaagctca gatctctaga gctgtcttgt cgccgcccag gattgacctg 2820
tgtgtaagtc ccaataaact cacctactca                                  2850
```

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<210> 10
<211> 383
<212> PRT
<213> Homo sapiens

<400> 10
Met Ser Ala Leu Phe Leu Gly Val Arg Val Arg Ala Glu Glu Ala Gly
1               5                   10                  15
Ala Arg Val Gln Gln Asn Val Pro Ser Gly Thr Asp Thr Gly Asp Pro
            20                  25                  30
Gln Ser Lys Pro Leu Gly Asp Trp Ala Ala Gly Thr Met Asp Pro Glu
        35                  40                  45
Ser Ser Ile Phe Ile Glu Asp Ala Ile Lys Tyr Phe Lys Glu Lys Val
    50                  55                  60
```

Ser Thr Gln Asn Leu Leu Leu Leu Leu Thr Asp Asn Glu Ala Trp Asn
65   70   75   80
Gly Phe Val Ala Ala Ala Glu Leu Pro Arg Asn Glu Ala Asp Glu Leu
                85   90   95
Arg Lys Ala Leu Asp Asn Leu Ala Arg Gln Met Ile Met Lys Asp Lys
            100   105   110
Asn Trp His Asp Lys Gly Gln Gln Tyr Arg Asn Trp Phe Leu Lys Glu
        115   120   125
Phe Pro Arg Leu Lys Ser Lys Leu Glu Asp Asn Ile Arg Arg Leu Arg
    130   135   140
Ala Leu Ala Asp Gly Val Gln Lys Val His Lys Gly Thr Thr Ile Ala
145   150   155   160
Asn Val Val Ser Gly Ser Leu Ser Ile Ser Ser Gly Ile Leu Thr Leu
                165   170   175
Val Gly Met Gly Leu Ala Pro Phe Thr Glu Gly Gly Ser Leu Val Leu
            180   185   190
Leu Glu Pro Gly Met Glu Leu Gly Ile Thr Ala Ala Leu Thr Gly Ile
        195   200   205
Thr Ser Ser Thr Ile Asp Tyr Gly Lys Lys Trp Trp Thr Gln Ala Gln
    210   215   220
Ala His Asp Leu Val Ile Lys Ser Leu Asp Lys Leu Lys Glu Val Lys
225   230   235   240
Glu Phe Leu Gly Glu Asn Ile Ser Asn Phe Leu Ser Leu Ala Gly Asn
                245   250   255
Thr Tyr Gln Leu Thr Arg Gly Ile Gly Lys Asp Ile Arg Ala Leu Arg
            260   265   270
Arg Ala Arg Ala Asn Leu Gln Ser Val Pro His Ala Ser Ala Ser Arg
        275   280   285
Pro Arg Val Thr Glu Pro Ile Ser Ala Glu Ser Gly Glu Gln Val Glu
    290   295   300
Arg Val Asn Glu Pro Ser Ile Leu Glu Met Ser Arg Gly Val Lys Leu
305   310   315   320
Thr Asp Val Ala Pro Val Ser Phe Phe Leu Val Leu Asp Val Val Tyr
                325   330   335
Leu Val Tyr Glu Ser Lys His Leu His Glu Gly Ala Lys Ser Glu Thr
            340   345   350
Ala Glu Glu Leu Lys Lys Val Ala Gln Glu Leu Glu Glu Lys Leu Asn
        355   360   365
Ile Leu Asn Asn Asn Tyr Lys Ile Leu Gln Ala Asp Gln Glu Leu
    370   375   380

<210> 11
<211> 3004
<212> DNA
<213> Homo sapiens

<400> 11
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tcagtctctg ccaggggaag attccttgac ttctggggtg atggagaaga aacaggctgt 180
gctgtgtccc taatgggaaa cgtggctgag acaggggagt gagaagggtg cgttgaagaa 240
tggtgcctgt ggcatgatgc cagctttgca atcatgagat tcaaaagcca cactgtggaa 300
ttgaggaggc cctgcagcga catggaggga gctgctttgc tgagtctctc tgtcctctgc 360
atctggatga gtgcactttt ccttggtgtg agagtgaggg cagaggaagc tggagcgagg 420
gtgcaacaaa acgttccaag tgggacagat actggagatc ctcaaagtaa gcccctcggt 480
gactgggctg ctggcaccat ggacccagag agcagtatct ttattgagga tgccattaag 540
tatttcaagg aaaaagtgag cacacagaat ctgctactcc tgctgactga taatgaggcc 600
tggaacggat tcgtggctgc tgctgaactg cccaggaatg aggcagatga gctccgtaaa 660

```
gctctggaca accttgcaag acaaatgatc atgaaagaca aaaactggca cgataaaggc 720
cagcagtaca gaaactggtt tctgaaagag tttcctcggt tgaaaagtaa gcttgaggat 780
aacataagaa ggctccgtgc ccttgcagat ggggttcaga aggtccacaa aggcaccacc 840
atcgccaatg tggtgtctgg ctctctcagc atttcctctg catcctgac cctcgtcggc 900
atgggtctgg caccettcac agagggaggc agccttgtac tcttggaacc tgggatggag 960
ttgggaatca cagccgcttt gaccgggatt accagcagta ccatagacta cggaaagaag 1020
tggtggacac aagcccaagc ccacgacctg gtcatcaaaa gccttgacaa attgaaggag 1080
gtgaaggagt ttttgggtga aacatatcc aactttcttt ccttagctgg caatacttac 1140
caactcacac gaggcattgg gaaggacatc cgtgccctca gacgagccag agccaatctt 1200
cagtcagtac cgcatgcctc agcctcacgc ccccgggtca ctgagccaat ctcagctgaa 1260
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aagctcacgg atgtggcccc tgtaagcttc tttcttgtgc tggatgtagt ctacctcgtg 1380
tacgaatcaa agcacttaca tgaggggca aagtcagaga cagctgagga gctgaagaag 1440
gtggctcagg agctggagga gaagctaaac attctcaaca ataattataa gattctgcag 1500
gcggaccaag aactgtgacc acagggcagg gcagccacca ggagagatat gcctggcagg 1560
ggccaggaca aaatgcaaac tttttttttt ttctgagaca gagtcttgct ctgtcgccaa 1620
gttggagtgc aatggtgcga tctcagctca ctgcaagctc tgcctcccgt gttcaagcga 1680
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aatttttgta tttttaatag agatggggtt tcaccatgtt ggccaggatg gtctcgatct 1800
cctgacctct tgatctgccc accttggcct cccaaagtgc tgggattaca ggcgtgagcc 1860
atcgcttttg acccaaatgc aaacatttta ttaggggat aaagagggtg aggtaaagtt 1920
tatggaactg agtgttaggg actttggcat ttccatagct gagcacagca ggggagggt 1980
taatgcagat ggcagtgcag caaggagaag gcaggaacat tggagcctgc aataagggaa 2040
aaatgggaac tggagagtgt ggggaatggg aagaagcagt ttactttaga ctaaagaata 2100
tattgggggg ccgggtgtag tggctcatgc ctgtaatccg agcactttgg gaggccaagg 2160
cgggcggatc acgaggtcag gagatcaaga ccatcctggc taacacagtg aaaccccgtc 2220
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tgggcggctg aggcaggaga atggcgtgaa cctgggaggt ggagcttgca gtgagccgag 2340
atatcgccac tgcactccag cctgggtgac agagcgagac tccatctcaa aaaaaaaaaa 2400
aaaagaata tattgacgga agaatagaga ggaggcttga aggaaccagc aatgagaagg 2460
ccaggaaaag aaagagctga aaatggagaa agcccaagag ttagaacagt tggatacagg 2520
agaagaaaca gcggctccac tacagaccca gccccaggtt caatgtcctc cgaagaatga 2580
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caggtggtgg gccatggcca tggtccccag ctgaggagca ggtgtccctg agaacccaaa 2820
cttcccagag agtatgtgag aaccaaccaa tgaaaacagt cccatcgctc ttacccggta 2880
agtaaacagt cagaaaatta gcatgaaagc agtttagcat gggaggaag ctcagatctc 2940
tagagctgtc ttgtcgccgc ccaggattga cctgtgtgta agtcccaata aactcaccta 3000
ctca                                                            3004
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<210> 12
<211> 414
<212> PRT
<213> Homo sapiens

<400> 12
Met Arg Phe Lys Ser His Thr Val Glu Leu Arg Arg Pro Cys Ser Asp
1               5                   10                  15
Met Glu Gly Ala Ala Leu Leu Arg Val Ser Val Leu Cys Ile Trp Met
            20                  25                  30
Ser Ala Leu Phe Leu Gly Val Arg Val Arg Ala Glu Glu Ala Gly Ala
        35                  40                  45
Arg Val Gln Gln Asn Val Pro Ser Gly Thr Asp Thr Gly Asp Pro Gln
    50                  55                  60
Ser Lys Pro Leu Gly Asp Trp Ala Ala Gly Thr Met Asp Pro Glu Ser
65                  70                  75                  80
Ser Ile Phe Ile Glu Asp Ala Ile Lys Tyr Phe Lys Glu Lys Val Ser
                85                  90                  95
```

```
Thr Gln Asn Leu Leu Leu Leu Leu Thr Asp Asn Glu Ala Trp Asn Gly
        100                 105                 110
Phe Val Ala Ala Ala Glu Leu Pro Arg Asn Glu Ala Asp Glu Leu Arg
        115                 120                 125
Lys Ala Leu Asp Asn Leu Ala Arg Gln Met Ile Met Lys Asp Lys Asn
        130                 135                 140
Trp His Asp Lys Gly Gln Gln Tyr Arg Asn Trp Phe Leu Lys Glu Phe
145                 150                 155                 160
Pro Arg Leu Lys Ser Lys Leu Glu Asp Asn Ile Arg Arg Leu Arg Ala
                165                 170                 175
Leu Ala Asp Gly Val Gln Lys Val His Lys Gly Thr Thr Ile Ala Asn
                180                 185                 190
Val Val Ser Gly Ser Leu Ser Ile Ser Ser Gly Ile Leu Thr Leu Val
                195                 200                 205
Gly Met Gly Leu Ala Pro Phe Thr Glu Gly Gly Ser Leu Val Leu Leu
        210                 215                 220
Glu Pro Gly Met Glu Leu Gly Ile Thr Ala Ala Leu Thr Gly Ile Thr
225                 230                 235                 240
Ser Ser Thr Ile Asp Tyr Gly Lys Lys Trp Trp Thr Gln Ala Gln Ala
                245                 250                 255
His Asp Leu Val Ile Lys Ser Leu Asp Lys Leu Lys Glu Val Lys Glu
                260                 265                 270
Phe Leu Gly Glu Asn Ile Ser Asn Phe Leu Ser Leu Ala Gly Asn Thr
        275                 280                 285
Tyr Gln Leu Thr Arg Gly Ile Gly Lys Asp Ile Arg Ala Leu Arg Arg
        290                 295                 300
Ala Arg Ala Asn Leu Gln Ser Val Pro His Ala Ser Ala Ser Arg Pro
305                 310                 315                 320
Arg Val Thr Glu Pro Ile Ser Ala Glu Ser Gly Glu Gln Val Glu Arg
                325                 330                 335
Val Asn Glu Pro Ser Ile Leu Glu Met Ser Arg Gly Val Lys Leu Thr
                340                 345                 350
Asp Val Ala Pro Val Ser Phe Phe Leu Val Leu Asp Val Val Tyr Leu
        355                 360                 365
Val Tyr Glu Ser Lys His Leu His Glu Gly Ala Lys Ser Glu Thr Ala
        370                 375                 380
Glu Glu Leu Lys Lys Val Ala Gln Glu Leu Glu Glu Lys Leu Asn Ile
385                 390                 395                 400
Leu Asn Asn Asn Tyr Lys Ile Leu Gln Ala Asp Gln Glu Leu
                405                 410
```

```
<210> 13
<211> 2298
<212> DNA
<213> Homo sapiens

<400> 13
ctaaaggtct ggttattatg cagatgcacg gctggaggtg ggatccacac agctcagaac 60
agctggatct tgctcacact ctttcaagag aagcttcctt ggacaaaagg accctgcctt 120
ggtgtgagag tgagggcaga gggagctgga gcaagtagaa tttctctaaa taccagctgg 180
ctggggccca ggagattaaa aaacaccggg ctaggttggt cttggcattt gctgacacgc 240
aaagggattg cagagatcca gcccctccaa cctccctctg tccacaggtg gctcacattc 300
agtcccacaa tttgctttct cctcctcaag ggttaagaaa aaaacgaac ccttccagtc 360
aggtcagtga ctggagagct ccaaggaaag tctctcagtg acctggctgc tggcaccatg 420
gactcagaaa agaaacgctt tactgaagag gccaccaaat acttccggga gagagtcagc 480
ccagtgcatc tgcaaatcct gctgactaac aatgaagcct ggaagagatt cgtgactgcg 540
gctgaattgc ccaggatgga ggcagatgct ctctacgaag ctctgaagaa gcttagaaca 600
tatgcagcta ttgaggacga atatgtgcag cagaaagatg agcagtttag ggaatggttt 660
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54

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ttgaaagagt ttccccaagt caagaggaag atccaggagt ccatagaaaa gcttcgtgcc 720
cttgcaaatg gtattgaaga ggtccacaga ggctgcacca tctccaatgt ggtgtccagc 780
tccactggcg ctgcctctgg catcatgtcc cttgctggtc ttgttttggc accatttaca 840
gcagggacga gtctggccct tactgcagct ggggtagggc tgggagcagc gtctgctgtg 900
actgggatca ccaccagcat cgtggagcac tcatacacat catcagcaga agctgaagcc 960
agcaggctga ctgcaaccag cattgaccga ttgaaggtat ttaaggaagt tatgcgtgac 1020
atcacaccca acttactttc ccttcttaat aattattacg aagccacaca aaccattggg 1080
agtgaaatcc gtgccatcag gcaagccaga gccagggccc gactcccgt gaccacctgg 1140
cgaatctcag ctggaagtgg tggtcaagca gagagaacga ttgcaggcac caccgggca 1200
gtgagcagag gagcccggat cctgagtgcg accacttcag gcatcttcct tgcactggat 1260
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gttgtagtgt gatggatccc ctttaggtta tttaggggta tatgtcccct gcttgaaccc 2040
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acccaaacat cctggagagt atgcgagaac ctaccaagaa aaacagtctc attactcata 2160
tacagcaggc aaagagacag aaaattaact gaaaagcagt ttagagactg ggggaggccg 2220
gatctctaga gccatcctgc tgagtgccct gtgtgtaagt cctaataaac tcacctactc 2280
accaaaaaaa aaaaaaaa                                                 2298
```

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<210> 14
<211> 331
<212> PRT
<213> Homo sapiens

<400> 14
Met Asp Ser Glu Lys Lys Arg Phe Thr Glu Glu Ala Thr Lys Tyr Phe
1               5                   10                  15
Arg Glu Arg Val Ser Pro Val His Leu Gln Ile Leu Leu Thr Asn Asn
            20                  25                  30
Glu Ala Trp Lys Arg Phe Val Thr Ala Ala Glu Leu Pro Arg Asp Glu
        35                  40                  45
Ala Asp Ala Leu Tyr Glu Ala Leu Lys Lys Leu Arg Thr Tyr Ala Ala
    50                  55                  60
Ile Glu Asp Glu Tyr Val Gln Gln Lys Asp Glu Gln Phe Arg Glu Trp
65                  70                  75                  80
Phe Leu Lys Glu Phe Pro Gln Val Lys Arg Lys Ile Gln Glu Ser Ile
                85                  90                  95
Glu Lys Leu Arg Ala Leu Ala Asn Gly Ile Glu Glu Val His Arg Gly
            100                 105                 110
Cys Thr Ile Ser Asn Val Val Ser Ser Thr Gly Ala Ala Ser Gly
            115                 120                 125
Ile Met Ser Leu Ala Gly Leu Val Leu Ala Pro Phe Thr Ala Gly Thr
    130                 135                 140
Ser Leu Ala Leu Thr Ala Ala Gly Val Gly Leu Gly Ala Ala Ser Ala
145                 150                 155                 160
Val Thr Gly Ile Thr Thr Ser Ile Val Glu His Ser Tyr Thr Ser Ser
                165                 170                 175
Ala Glu Ala Glu Ala Ser Arg Leu Thr Ala Thr Ser Ile Asp Arg Leu
            180                 185                 190
```

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Lys Val Phe Lys Glu Val Met Arg Asp Ile Thr Pro Asn Leu Leu Ser
         195                 200                 205
Leu Leu Asn Asn Tyr Tyr Glu Ala Thr Gln Thr Ile Gly Ser Glu Ile
         210                 . 215                220
Arg Ala Ile Arg Gln Ala Arg Ala Arg Ala Arg Leu Pro Val Thr Thr
225                 230                 235                 240
Trp Arg Ile Ser Ala Gly Ser Gly Gly Gln Ala Glu Arg Thr Ile Ala
             245                 250                 255
Gly Thr Thr Arg Ala Val Ser Arg Gly Ala Arg Ile Leu Ser Ala Thr
         260                 265                 270
Thr Ser Gly Ile Phe Leu Ala Leu Asp Val Val Asn Leu Val Tyr Glu
         275                 280                 285
Ser Lys His Leu His Glu Gly Ala Lys Ser Ala Ser Ala Glu Glu Leu
         290                 295                 300
Arg Arg Gln Ala Gln Glu Leu Glu Glu Asn Leu Met Glu Leu Thr Gln
305                 310                 315                 320
Ile Tyr Gln Arg Leu Asn Pro Cys His Thr His
                 325                 330
```

<210> 15
<211> 1316
<212> DNA
<213> Homo sapiens

<400> 15
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cggggcgcac ggcaccgagg agagcgcggg agccaacctg ggcgcatcat gcgcagggcc 120
cgggacgctg ggccggtcta caccgccgcc tgggtcacgt ggccccggacg ggccggcggc 180
tgccccggcc gggggggcggg ggtcgcgccg gggttgcgct ggacgacgga gagcggcggg 240
cccgcagcgg cctggagcct cccaacccgc gccgcgctgg ccctcgagcg taggagccgc 300
cccctgcccc cccgcgccgg ccccgcgccc ggccgcccgc ccctatata gcgcgccccca 360
gcagggcccg cgccaggccg ccagcctcgg agtgggcgcg ggacagtgcg cggcgccccg 420
cagccaggcc cccgccccccg ccgcatccac ctcctccgcc gcctgcgacc caacgggcgc 480
cccccgccgg cagctcgcgc cgggcccccg cggccaccat gaagaaggag gtgtgctccg 540
tggccttcct caaggccgtg ttcgcagagt tcttggccac cctcatcttc gtcttctttg 600
gcctgggctc ggccctcaag tggccgtcgg cgctgcctac catcctgcag atcgcgctgg 660
cgtttggcct ggccataggc acgctggccc aggccctggg accgtgagc ggcggccaca 720
tcaaccccgc catcaccctg gccctcttgg tgggcaacca gatctcgctg ctccgggctt 780
tcttctacgt ggcggcccag ctggtgggcg ccattgccgg ggctggcatc ctctacggtg 840
tggcaccgct caatgcccgg ggcaatctgg ccgtcaacgc gctcaacaac aacacaacgc 900
agggccaggc catggtggtg gagctgattc tgaccttcca gctggcactc tgcatcttcg 960
cctccactga ctcccgccgc accagccctg tgggctcccc agccctgtcc attggcctgt 1020
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gctcttttgg ccctgcggtg gtcatgaatc ggttcagccc cgctcactgg gttttctggg 1140
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actccctgag cctgagtgag cgtgtggcca tcatcaaagg cacgtatgag cctgacgagg 1260
actgggagga gcagcgggaa gagcggaaga agaccatgga gctgaccacc cgctga 1316
```

<210> 16
<211> 265
<212> PRT
<213> Homo sapiens

<400> 16
```
Met Lys Lys Glu Val Cys Ser Val Ala Phe Leu Lys Ala Val Phe Ala
 1               5                  10                  15
Glu Phe Leu Ala Thr Leu Ile Phe Val Phe Phe Gly Leu Gly Ser Ala
             20                  25                  30
```

```
Leu Lys Trp Pro Ser Ala Leu Pro Thr Ile Leu Gln Ile Ala Leu Ala
        35                  40                  45
Phe Gly Leu Ala Ile Gly Thr Leu Ala Gln Ala Leu Gly Pro Val Ser
    50                  55                  60
Gly Gly His Ile Asn Pro Ala Ile Thr Leu Ala Leu Leu Val Gly Asn
65                  70                  75                  80
Gln Ile Ser Leu Leu Arg Ala Phe Phe Tyr Val Ala Ala Gln Leu Val
                85                  90                  95
Gly Ala Ile Ala Gly Ala Gly Ile Leu Tyr Gly Val Ala Pro Leu Asn
            100                 105                 110
Ala Arg Gly Asn Leu Ala Val Asn Ala Leu Asn Asn Asn Thr Thr Gln
        115                 120                 125
Gly Gln Ala Met Val Val Glu Leu Ile Leu Thr Phe Gln Leu Ala Leu
    130                 135                 140
Cys Ile Phe Ala Ser Thr Asp Ser Arg Arg Thr Ser Pro Val Gly Ser
145                 150                 155                 160
Pro Ala Leu Ser Ile Gly Leu Ser Val Thr Leu Gly His Leu Val Gly
            165                 170                 175
Ile Tyr Phe Thr Gly Cys Ser Met Asn Pro Ala Arg Ser Phe Gly Pro
        180                 185                 190
Ala Val Val Met Asn Arg Phe Ser Pro Ala His Trp Val Phe Trp Val
    195                 200                 205
Gly Pro Ile Val Gly Ala Val Leu Ala Ala Ile Leu Tyr Phe Tyr Leu
    210                 215                 220
Leu Phe Pro Asn Ser Leu Ser Leu Ser Glu Arg Val Ala Ile Ile Lys
225                 230                 235                 240
Gly Thr Tyr Glu Pro Asp Glu Asp Trp Glu Glu Gln Arg Glu Glu Arg
            245                 250                 255
Lys Lys Thr Met Glu Leu Thr Thr Arg
            260                 265
```

```
<210> 17
<211> 1258
<212> DNA
<213> Homo sapiens

<400> 17
cacatatata atgaaaagta atcagtctcc aaagttttta tgtgtcatgt aagattactg 60
cttgcctctc taaggaaggt cgtgactgtt taaatagacg ggcaaggtgg aacctttttga 120
aagatgagct tttgaatata agttgtctgc tagatcatgg tttgtattga actaacaagg 180
tttgcagatc tgctgactta tataaagctt tttgattcct actaagcttt aagatttaaa 240
aaatgttcaa tgttgaaatt tctgtggggc tctatttttg ctttggcttt ctggtgagag 300
agtgaggaag cattctttcc ttcactaagt ttgtctttct tgtcttctgg atagattgat 360
tttaagagac taagggaatt tacaaactaa agatttagt catctggtgg aaaaggagac 420
tttaagattg tttagggctg ggcggggtga ctcacatctg taatcccagc actttgggag 480
gccgaggcag gcagaacact tgaaggagtt caagaccagc gtggccaacg tggtgaaacc 540
ctgtctctac taaaaataca aaaattgttt agctctgttt ttcataatag aaatagaaaa 600
ggtaaaattg ctttctttct gaaaagaaca agtattgttc atccaagaag ggttttgtg 660
actgaatcag cagtgcctgc cctagtcata gctgtgcttc aaaaacctca gcatgattag 720
tgttggagca aaacaaggaa gcaaagcaaa tactgttttt gaaattctat ctgttgcttg 780
aactattttg taataattaa actttgatgt tgagaaatca caactttatt gtacacttca 840
ttgcaacttg aaattcatgg tcttaaagtg agatttgaat ttctattgag cgcctttaaa 900
aaagtaatac caaaccataa agttaaaatc tatgtatatt gagtcatatc taaaaccacg 960
tataaacata aattgtattt cctgttttaa ttccagggga agtactgttt gggaaagcta 1020
ttattaggta aatgttttac aaattactgt ttctcacttt cagtcatacc ctaatgatcc 1080
cagcaagata atgtcctgtc ttctaagatg tgcatcaagc ctggtacata ctgaaaaccc 1140
tataaggtcc tggataattt ttgtttgatt attcattgaa gaaacatttta ttttccaatt 1200
gtgtgaagtt tttgactgtt aataaagaa tctgtcaacc atcaaaaaaa aaaaaaaa 1258
```

```
<210> 18
<211> 22
<212> PRT
<213> Homo sapiens

<400> 18
Met Val Cys Ile Glu Leu Thr Arg Phe Ala Asp Leu Leu Thr Tyr Ile
1               5                   10                  15
Lys Leu Phe Asp Ser Tyr
            20
```

```
<210> 19
<211> 983
<212> DNA
<213> Homo sapiens

<400> 19
gtggaattca tggcatctac ttcgtatgac tattgcagag tgcccatgga agacgggat   60
aagcgctgta agcttctgct ggggatagga attctggtgc tcctgatcat cgtgattctg  120
ggggtgccct tgattatctt caccatcaag gccaacagcg aggcctgccg ggacggcctt  180
cgggcagtga tggagtgtcg caatgtcacc atctcctgc aacaagagct gaccgaggcc   240
cagaagggct tcaggatgt ggaggcccag gccgccacct gcaaccacac tgtgatggcc   300
ctaatggctt ccctggatgc agagaaggcc caaggacaaa agaaagtgga ggagcttgag  360
ggagagatca ctacattaaa ccataagctt caggacgcgt ctgcagaggt ggagcgactg  420
agaagagaaa accaggtctt aagcgtgaga atcgcggaca gaagtacta ccccagctcc   480
caggactcca gctccgctgc ggcgccccag ctgctgattg tgctgctggg cctcagcgct  540
ctgctgcagt gagatcccag gaagctggca catcttggaa ggtccgtcct gctcggcttt  600
tcgcttgaac attcccttga tctcatcagt tctgagcggg tcatgggca acacggttag   660
cggggagagc acggggtagc cggagaaggg cctctggagc aggtctggag gggccatggg  720
gcagtcctgg gtgtggggac acagtcgggt tgacccaggg ctgtctccct ccagagcctc  780
cctccggaca atgagtcccc cctcttgtct cccaccctga gattgggcat ggggtgcggt  840
gtggggggca tgtgctgcct gttgttatgg gtttttttg cgggggggt tgctttttc    900
tggggtcttt gagctccaaa aaataaacac ttcctttgag ggagagcaaa aaaaaaaaa   960
aaaaaaaaaa aaaaaaaaa aaa                                           983
```

```
<210> 20
<211> 180
<212> PRT
<213> Homo sapiens

<400> 20
Met Ala Ser Thr Ser Tyr Asp Tyr Cys Arg Val Pro Met Glu Asp Gly
1               5                   10                  15
Asp Lys Arg Cys Lys Leu Leu Leu Gly Ile Gly Ile Leu Val Leu Leu
            20                  25                  30
Ile Ile Val Ile Leu Gly Val Pro Leu Ile Ile Phe Thr Ile Lys Ala
            35                  40                  45
Asn Ser Glu Ala Cys Arg Asp Gly Leu Arg Ala Val Met Glu Cys Arg
        50                  55                  60
Asn Val Thr His Leu Leu Gln Gln Glu Leu Thr Glu Ala Gln Lys Gly
65                  70                  75                  80
Phe Gln Asp Val Glu Ala Gln Ala Ala Thr Cys Asn His Thr Val Met
            85                  90                  95
Ala Leu Met Ala Ser Leu Asp Ala Glu Lys Ala Gln Gly Gln Lys Lys
            100                 105                 110
Val Glu Glu Leu Glu Gly Glu Ile Thr Thr Leu Asn His Lys Leu Gln
            115                 120                 125
```

```
Asp Ala Ser Ala Glu Val Glu Arg Leu Arg Arg Glu Asn Gln Val Leu
    130             135             140
Ser Val Arg Ile Ala Asp Lys Lys Tyr Tyr Pro Ser Ser Gln Asp Ser
145             150             155             160
Ser Ser Ala Ala Ala Pro Gln Leu Leu Ile Val Leu Leu Gly Leu Ser
                165             170             175
Ala Leu Leu Gln
        180
```

```
<210> 21
<211> 4859
<212> DNA
<213> Homo sapiens

<400> 21
cacgttgggt gacataatgg ggttttttta attatagatt cacactgcat ttattcatca 60
cccctgtcct ctcatccata actcaaattt actaccagca acacaaaata caaagatgtg 120
tccagtttca ctacagctct tcgcgtttac aagtgtcgag cgcttgcttt cggaacgccc 180
ttgtgattgg ccgagccaat gccagtgaca tcaaccaact tacttttgat tggaaggctg 240
gttgctggga ctgtagcgtt tgcaggaagt cacttaactg tttgggagct ggaaaaccga 300
agctgaagtt ctcttttgcc ataggaacga gcgcaactga ctaggaaaga tgtgtcccaa 360
agctccgcaa gctggaacgt gagccaggag gcccggaccg gccacgggac cgcgaggcac 420
tccgaaagtg tgcggctgcc ccttccctgc ctcccagctg ttaccctttt aaatgtcagt 480
gttcgaggct gtaggggtag cacgaggcag cgaaacggaa cagtcggatt ggccgcacgc 540
ctcagttcta gacgcacctc tccaccgaag ccgttctgac tggcagggg agaaagtaaa 600
cagagttgaa tcaccctccc cactggccaa ttggaggggg tttggtttgt gacgtgatgg 660
gattctgcga aattgttact gagcaagaga atgccggaac gtgcggaccg gccggagcag 720
gggttcagaa gccgtcagtg gactcgggaa aaagtgtctc ttagacctgg cgctcggcgg 780
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aaagatgagg ggctgaggtc gcctcggcgc tcctgcgagt cggaagcgcc ccgcgccccc 1140
gcccccttgg ccgccgcgcc gtgccgggcg ggcgggtcgt cgtccgaggc cagggagggc 1200
gagccgaacc tccgcagcca ccgccaagtt tgtccgcgcc gcctgggctg ccgtcgcccg 1260
caccatgtcc gcggccgcct acatggactt cgtggctgcc cagtgtctgg tttccatttc 1320
gaaccgcgct gcggtgccgg agcatggggt cgctccggac gccgagcggc tgcgactacc 1380
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cgcgccagc ccgctctccc tcctccatcc tggagtggct gcgaagggga aacacgcctc 1680
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aaagcacgcc cggcggcaca ccgagttcca ccccagcatg atcaagcgat cgaaaaaggc 1980
gctggccaac gctttgtgag gtgctgcccg tggaagccag ggagggatgg accccgaaag 2040
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caactctctg aacccgtctc tgactgttgg agggaaggca aatgcttttg ggttttttgg 2280
tttttgtttt tgttttttttt tctccttta ttttttttgcg ggggagggta gggagtgggt 2340
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atgtccgctg ggtgagttga cctgtggcct cgcacagtga ttctaggccc tttatgcttg 2460
ctgtctctca gaattgtttt cttacctttt aatgtaatga cgagtgtgct tcagtttgtt 2520
tagcaaaacc actctcttga atcacgttaa cttttgagat taaaaaaaaa aacgccatag 2580
```

```
cacagctgtc tttatgcaag caagagcaca tctactccag catgatctgt catctaaaga 2640
cttgaaaaca aaaaacagtt acttatagtc aatgggtaag cagagtctga atttatacta 2700
atcaagacaa acctttgaaa ggttacacta agtacagaac ttttaaacct tgctttgtat 2760
gagttgtact ttttgaacat aagctgcact tttattttct aatgcagagg atgaataagt 2820
taaatacatg ctttgaggat agaagcagat gttctgtttg gcaccacgtt ataatctgct 2880
tattttacaa tatacacgtt tccctaagaa atcatgcgca gagatgtgag ggcagaatat 2940
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aacagaattt taactctatt aacttttcca aattttccta tgctttttagt taacatcatt 3060
attgtatcct aatgccacta ggggagagag cttttgactc tgttgggttt tatttgaatg 3120
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gtccaaaata agttttaagg acatctgttc ttccatacga aataggttag gctgcctatt 4080
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aaaaaatgca catttacttc ccaaaaaagt tgttacttgc cttttcaagt gtgacaaact 4380
cacatttgat attctcttat atgttatagt aatgtaacgt ataaactcaa gccttttttat 4440
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tgaaacaatt acttttttata ttaaaaagac catggattta acttatgaaa atccaaatgc 4620
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tgtttaaaaa gaaagaaaag ttgttttatc taatactgta agtagttgtc atattctgga 4740
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```

```
<210> 22
<211> 244
<212> PRT
<213> Homo sapiens

<400> 22
Met Ser Ala Ala Ala Tyr Met Asp Phe Val Ala Ala Gln Cys Leu Val
1               5                   10                  15
Ser Ile Ser Asn Arg Ala Ala Val Pro Glu His Gly Val Ala Pro Asp
            20                  25                  30
Ala Glu Arg Leu Arg Leu Pro Glu Arg Glu Val Thr Lys Glu His Gly
        35                  40                  45
Asp Pro Gly Asp Thr Trp Lys Asp Tyr Cys Thr Leu Val Thr Ile Ala
    50                  55                  60
Lys Ser Leu Leu Asp Leu Asn Lys Tyr Arg Pro Ile Gln Thr Pro Ser
65                  70                  75                  80
Val Cys Ser Asp Ser Leu Glu Ser Pro Asp Glu Asp Met Gly Ser Asp
                85                  90                  95
Ser Asp Val Thr Thr Glu Ser Gly Ser Ser Pro Ser His Ser Pro Glu
            100                 105                 110
```

```
Glu Arg Gln Asp Pro Gly Ser Ala Pro Ser Pro Leu Ser Leu Leu His
        115                 120                 125
Pro Gly Val Ala Ala Lys Gly Lys His Ala Ser Glu Lys Arg His Lys
        130                 135                 140
Cys Pro Tyr Ser Gly Cys Gly Lys Val Tyr Gly Lys Ser Ser His Leu
145                 150                 155                 160
Lys Ala His Tyr Arg Val His Thr Gly Glu Arg Pro Phe Pro Cys Thr
                165                 170                 175
Trp Pro Asp Cys Leu Lys Lys Phe Ser Arg Ser Asp Glu Leu Thr Arg
            180                 185                 190
His Tyr Arg Thr His Thr Gly Glu Lys Gln Phe Arg Cys Pro Leu Cys
        195                 200                 205
Glu Lys Arg Phe Met Arg Ser Asp His Leu Thr Lys His Ala Arg Arg
        210                 215                 220
His Thr Glu Phe His Pro Ser Met Ile Lys Arg Ser Lys Lys Ala Leu
225                 230                 235                 240
Ala Asn Ala Leu
```

```
<210> 23
<211> 1304
<212> DNA
<213> Homo sapiens

<400> 23
ttcccagatg cacaggagga gaagcaggag ctgtcgggaa gatcagaagc cagtcatgga 60
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ggccccggag agcaagtgca gccgcggagc cctgtacaca ggcttttcca tcctggtgac 180
tctgctcctc gctggccagg ccaccaccgc ctacttcctg taccagcagc agggccggct 240
ggacaaactg acagtcacct cccagaacct gcagctggag aacctgcgca tgaagcttcc 300
caagcctccc aagcctgtga gcaagatgcg catggccacc ccgctgctga tgcaggcgct 360
gcccatggga gccctgcccc aggggcccat gcagaatgcc accaagtatg caacatgac 420
agaggaccat gtgatgcacc tgctccagaa tgctgacccc ctgaaggtgt acccgccact 480
gaaggggagc ttcccggaga acctgagaca ccttaagaac accatggaga ccatagactg 540
gaaggtcttt gagagctgga tgcaccattg gctcctgttt gaaatgagca ggcactcctt 600
ggagcaaaag cccactgacg ctccaccgaa agagtcactg gaactggagg acccgtcttc 660
tgggctgggt gtgaccaagc aggatctggg cccagtcccc atgtgagagc agcagaggcg 720
gtcttcaaca tcctgccagc cccacacagc tacagctttc ttgctccctt cagcccccag 780
cccctccccc atgtcccacc ctgtacctca tcccatgaga cctggtgcct ggctctttcg 840
tcacccttgt acaagacaaa ccaagtcgga acagcagata acaatgcagc aaggccctgc 900
tgcccaatct ccatctgtca acaggggcgt gaggtcccag gaagtggcca aaagctagac 960
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atgctgactg tcctctcccc tccagccttt ggccttggct tttctagcct atttacctgc 1140
aggctgagcc actctcttcc ctttccccag catcactccc caaggaagag ccaatgtttt 1200
ggacccataa tcctttctgc cgacccctag ttccctctgc tcagccaagc ttgttatcag 1260
ctttcagggc catggttcac attagaataa aaggtagtaa ttag      1304
```

```
<210> 24
<211> 232
<212> PRT
<213> Homo sapiens

<400> 24
Met His Arg Arg Arg Ser Arg Ser Cys Arg Glu Asp Gln Lys Pro Val
1               5               10              15
Met Asp Asp Gln Arg Asp Leu Ile Ser Asn Asn Glu Gln Leu Pro Met
            20              25              30
```

```
Leu Gly Arg Arg Pro Gly Ala Pro Glu Ser Lys Cys Ser Arg Gly Ala
        35              40              45
Leu Tyr Thr Gly Phe Ser Ile Leu Val Thr Leu Leu Leu Ala Gly Gln
    50              55                      60
Ala Thr Thr Ala Tyr Phe Leu Tyr Gln Gln Gln Gly Arg Leu Asp Lys
65              70              75                      80
Leu Thr Val Thr Ser Gln Asn Leu Gln Leu Glu Asn Leu Arg Met Lys
            85              90                      95
Leu Pro Lys Pro Pro Lys Pro Val Ser Lys Met Arg Met Ala Thr Pro
        100             105             110
Leu Leu Met Gln Ala Leu Pro Met Gly Ala Leu Pro Gln Gly Pro Met
        115             120             125
Gln Asn Ala Thr Lys Tyr Gly Asn Met Thr Glu Asp His Val Met His
    130             135             140
Leu Leu Gln Asn Ala Asp Pro Leu Lys Val Tyr Pro Pro Leu Lys Gly
145             150             155             160
Ser Phe Pro Glu Asn Leu Arg His Leu Lys Asn Thr Met Glu Thr Ile
            165             170             175
Asp Trp Lys Val Phe Glu Ser Trp Met His His Trp Leu Leu Phe Glu
        180             185             190
Met Ser Arg His Ser Leu Glu Gln Lys Pro Thr Asp Ala Pro Pro Lys
        195             200             205
Glu Ser Leu Glu Leu Glu Asp Pro Ser Ser Gly Leu Gly Val Thr Lys
    210             215             220
Gln Asp Leu Gly Pro Val Pro Met
225             230
```

```
<210> 25
<211> 1615
<212> DNA
<213> Homo sapiens

<400> 25
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<210> 26
<211> 499
<212> PRT
<213> Homo sapiens

<400> 26

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            20                  25                  30
Lys Glu Ala Ala Gly Pro Ala Pro Ser Pro Met Arg Ala Ala Asn Arg
        35                  40                  45
Ser His Ser Ala Gly Arg Thr Pro Gly Arg Thr Pro Gly Lys Ser Ser
    50                  55                  60
Ser Lys Val Gln Thr Thr Pro Ser Lys Pro Gly Gly Asp Arg Tyr Ile
65                  70                  75                  80
Pro His Arg Ser Ala Ala Gln Met Glu Val Ala Ser Phe Leu Leu Ser
                85                  90                  95
Lys Glu Asn Gln Pro Glu Asn Ser Gln Thr Pro Thr Lys Lys Glu His
            100                 105                 110
Gln Lys Ala Trp Ala Leu Asn Leu Asn Gly Phe Asp Val Glu Glu Ala
        115                 120                 125
Lys Ile Leu Arg Leu Ser Gly Lys Pro Gln Asn Ala Pro Glu Gly Tyr
    130                 135                 140
Gln Asn Arg Leu Lys Val Leu Tyr Ser Gln Lys Ala Thr Pro Gly Ser
145                 150                 155                 160
Ser Arg Lys Thr Cys Arg Tyr Ile Pro Ser Leu Pro Asp Arg Ile Leu
                165                 170                 175
Asp Ala Pro Glu Ile Arg Asn Asp Tyr Tyr Leu Asn Leu Val Asp Trp
            180                 185                 190
Ser Ser Gly Asn Val Leu Ala Val Ala Leu Asp Asn Ser Val Tyr Leu
        195                 200                 205
Trp Ser Ala Ser Ser Gly Asp Ile Leu Gln Leu Leu Gln Met Glu Gln
    210                 215                 220
Pro Gly Glu Tyr Ile Ser Ser Val Ala Trp Ile Lys Glu Gly Asn Tyr
225                 230                 235                 240
Leu Ala Val Gly Thr Ser Ser Ala Glu Val Gln Leu Trp Asp Val Gln
                245                 250                 255
Gln Gln Lys Arg Leu Arg Asn Met Thr Ser His Ser Ala Arg Val Gly
            260                 265                 270
Ser Leu Ser Trp Asn Ser Tyr Ile Leu Ser Ser Gly Ser Arg Ser Gly
        275                 280                 285
His Ile His His His Asp Val Arg Val Ala Glu His His Val Ala Thr
    290                 295                 300
Leu Ser Gly His Ser Gln Glu Val Cys Gly Leu Arg Trp Ala Pro Asp
305                 310                 315                 320
Gly Arg His Leu Ala Ser Gly Gly Asn Asp Asn Leu Val Asn Val Trp
                325                 330                 335
Pro Ser Ala Pro Gly Glu Gly Gly Trp Val Pro Leu Gln Thr Phe Thr
            340                 345                 350
Gln His Gln Gly Ala Val Lys Ala Val Ala Trp Cys Pro Trp Gln Ser
        355                 360                 365
Asn Val Leu Ala Thr Gly Gly Gly Thr Ser Asp Arg His Ile Arg Ile
    370                 375                 380
Trp Asn Val Cys Ser Gly Ala Cys Leu Ser Ala Val Asp Ala His Ser
385                 390                 395                 400
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Gln Val Cys Ser Ile Leu Trp Ser Pro His Tyr Lys Glu Leu Ile Ser
            405                 410                 415
Gly His Gly Phe Ala Gln Asn Gln Leu Val Ile Trp Lys Tyr Pro Thr
            420                 425                 430
Met Ala Lys Val Ala Glu Leu Lys Gly His Thr Ser Arg Val Leu Ser
            435                 440                 445
Leu Thr Met Ser Pro Asp Gly Ala Thr Val Ala Ser Ala Ala Ala Asp
        450                 455                 460
Glu Thr Leu Arg Leu Trp Arg Cys Phe Glu Leu Asp Pro Ala Arg Arg
465                 470                 475                 480
Arg Glu Arg Glu Lys Ala Ser Ala Ala Lys Ser Ser Leu Ile His Gln
                485                 490                 495
Gly Ile Arg
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<210> 27
<211> 2103
<212> DNA
<213> Homo sapiens

<400> 27
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<210> 28
<211> 168
<212> PRT
<213> Homo sapiens

<400> 28
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Asp Leu Glu Gln Leu Thr Ser Leu Leu Gln Asn Asn Val Asn Val Asn
            20                  25                  30
Ala Gln Asn Gly Phe Gly Arg Thr Ala Leu Gln Val Met Lys Leu Gly
        35                  40                  45
Asn Pro Glu Ile Ala Arg Arg Leu Leu Leu Arg Gly Ala Asn Pro Asp
    50                  55                  60
Leu Lys Asp Arg Thr Gly Phe Ala Val Ile His Asp Ala Ala Arg Ala
65                  70                  75                  80
Gly Phe Leu Asp Thr Leu Gln Thr Leu Leu Glu Phe Gln Ala Asp Val
                85                  90                  95
Asn Ile Glu Asp Asn Glu Gly Asn Leu Pro Leu His Leu Ala Ala Lys
                100                 105                 110
Glu Gly His Leu Arg Val Val Glu Phe Leu Val Lys His Thr Ala Ser
            115                 120                 125
Asn Val Gly His Arg Asn His Lys Gly Asp Thr Ala Cys Asp Leu Ala
    130                 135                 140
Arg Leu Tyr Gly Arg Asn Glu Val Val Ser Leu Met Gln Ala Asn Gly
145                 150                 155                 160
Ala Gly Gly Ala Thr Asn Leu Gln
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<210> 29
<211> 4049
<212> DNA
<213> Homo sapiens

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<210> 30
<211> 184
<212> PRT
<213> Homo sapiens

<400> 30

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Thr Leu Leu Pro Ala Ala Glu Gly Lys Lys Lys Gly Ser Gln Gly Ala
            20                  25                  30
Ile Pro Pro Pro Asp Lys Ala Gln His Asn Asp Ser Glu Gln Thr Gln
        35                  40                  45
Ser Pro Gln Gln Pro Gly Ser Arg Asn Arg Gly Arg Gly Gln Gly Arg
```

```
        50                      55                      60
Gly Thr Ala Met Pro Gly Glu Glu Val Leu Glu Ser Ser Gln Glu Ala
65                      70                      75                      80
Leu His Val Thr Glu Arg Lys Tyr Leu Lys Arg Asp Trp Cys Lys Thr
                    85                      90                      95
Gln Pro Leu Lys Gln Thr Ile His Glu Glu Gly Cys Asn Ser Arg Thr
                100                     105                     110
Ile Ile Asn Arg Phe Cys Tyr Gly Gln Cys Asn Ser Phe Tyr Ile Pro
            115                     120                     125
Arg His Ile Arg Lys Glu Glu Gly Ser Phe Gln Ser Cys Ser Phe Cys
        130                     135                     140
Lys Pro Lys Lys Phe Thr Thr Met Met Val Thr Leu Asn Cys Pro Glu
145                     150                     155                     160
Leu Gln Pro Pro Thr Lys Lys Lys Arg Val Thr Arg Val Lys Gln Cys
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Arg Cys Ile Ser Ile Asp Leu Asp
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<210> 31
<211> 3443
<212> DNA
<213> Homo sapiens

<400> 31

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<213> Homo sapiens

<400> 32

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Asp Leu Gln Asn Leu Ala Pro Gly Thr His Pro Pro Phe Ile Thr Phe
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Asn Ser Glu Val Lys Thr Asp Val Asn Lys Ile Glu Glu Phe Leu Glu
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Glu Ser Asn Thr Ala Gly Met Asp Ile Phe Ala Lys Phe Ser Ala Tyr
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Ile Lys Asn Ser Arg Pro Glu Ala Asn Glu Ala Leu Glu Arg Gly Leu
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Leu Lys Thr Leu Gln Lys Leu Asp Glu Tyr Leu Asn Ser Pro Leu Pro
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<400> 36

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Asp Arg Asp Val Trp Lys Pro Glu Pro Cys Arg Ile Cys Val Cys Asp
    50                  55                  60
Asn Gly Lys Val Leu Cys Asp Asp Val Ile Cys Asp Glu Thr Lys Asn
65                  70                  75                  80
Cys Pro Gly Ala Glu Val Pro Glu Gly Glu Cys Cys Pro Val Cys Pro
                85                  90                  95
Asp Gly Ser Glu Ser Pro Thr Asp Gln Glu Thr Thr Gly Val Glu Gly
            100                 105                 110
Pro Lys Gly Asp Thr Gly Pro Arg Gly Pro Arg Gly Pro Ala Gly Pro
            115                 120                 125
Pro Gly Arg Asp Gly Ile Pro Gly Gln Pro Gly Leu Pro Gly Pro Pro
.    130                 135                 140
Gly Pro Pro Gly Pro Pro Gly Pro Pro Gly Leu Gly Gly Asn Phe Ala
145                 150                 155                 160
Pro Gln Leu Ser Tyr Gly Tyr Asp Glu Lys Ser Thr Gly Gly Ile Ser
                165                 170                 175
Val Pro Gly Pro Met Gly Pro Ser Gly Pro Arg Gly Leu Pro Gly Pro
            180                 185                 190
Pro Gly Ala Pro Gly Pro Gln Gly Phe Gln Gly Pro Pro Gly Glu Pro
    .195                 200                 .   205
Gly Glu Pro Gly Ala Ser Gly Pro Met Gly Pro Arg Gly Pro Pro Gly
    210                 215                 220
Pro Pro Gly Lys Asn Gly Asp Asp Gly Glu Ala Gly Lys Pro Gly Arg
225                 230                 235                 240
Pro Gly Glu Arg Gly Pro Pro Gly Pro Gln Gly Ala Arg Gly Leu Pro
                245                 250                 255
Gly Thr Ala Gly Leu Pro Gly Met Lys Gly His Arg Gly Phe Ser Gly
            260                 265                 270
Leu Asp Gly Ala Lys Gly Asp Ala Gly Pro Ala Gly Pro Lys Gly Glu
            275                 280                 285
Pro Gly Ser Pro Gly Glu Asn Gly Ala Pro Gly Gln Met Gly Pro Arg
            290                 295                 300
Gly Leu Pro Gly Glu Arg Gly Arg Pro Gly Ala Pro Gly Pro Ala Gly
305                 310                 315                 320
Ala Arg Gly Asn Asp Gly Ala Thr Gly Ala Ala Gly Pro Pro Gly Pro
                325                 330                 335
Thr Gly Pro Ala Gly Pro Pro Gly Phe Pro Gly Ala Val Gly Ala Lys
            340                 345                 350
Gly Glu Ala Gly Pro Gln Gly Pro Arg Gly Ser Glu Gly Pro Gln Gly
            355                 360                 365
Val Arg Gly Glu Pro Gly Pro Pro Gly Pro Ala Gly Ala Ala Gly Pro
    370                 375                 380
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Ala Gly Asn Pro Gly Ala Asp Gly Gln Pro Gly Ala Lys Gly Ala Asn
385             390             395             400
Gly Ala Pro Gly Ile Ala Gly Ala Pro Gly Phe Pro Gly Ala Arg Gly
            405             410             415
Pro Ser Gly Pro Gln Gly Pro Gly Gly Pro Pro Gly Pro Lys Gly Asn
        420             425             430
Ser Gly Glu Pro Gly Ala Pro Gly Ser Lys Gly Asp Thr Gly Ala Lys
        435             440             445
Gly Glu Pro Gly Pro Val Gly Val Gln Gly Pro Pro Gly Pro Ala Gly
    450             455             460
Glu Glu Gly Lys Arg Gly Ala Arg Gly Glu Pro Gly Pro Thr Gly Leu
465             470             475             480
Pro Gly Pro Pro Gly Glu Arg Gly Gly Pro Gly Ser Arg Gly Phe Pro
            485             490             495
Gly Ala Asp Gly Val Ala Gly Pro Lys Gly Pro Ala Gly Glu Arg Gly
        500             505             510
Ser Pro Gly Pro Ala Gly Pro Lys Gly Ser Pro Gly Glu Ala Gly Arg
        515             520             525
Pro Gly Glu Ala Gly Leu Pro Gly Ala Lys Gly Leu Thr Gly Ser Pro
    530             535             540
Gly Ser Pro Gly Pro Asp Gly Lys Thr Gly Pro Pro Gly Pro Ala Gly
545             550             555             560
Gln Asp Gly Arg Pro Gly Pro Pro Gly Pro Pro Gly Ala Arg Gly Gln
            565             570             575
Ala Gly Val Met Gly Phe Pro Gly Pro Lys Gly Ala Ala Gly Glu Pro
            580             585             590
Gly Lys Ala Gly Glu Arg Gly Val Pro Gly Pro Pro Gly Ala Val Gly
        595             600             605
Pro Ala Gly Lys Asp Gly Glu Ala Gly Ala Gln Gly Pro Pro Gly Pro
    610             615             620
Ala Gly Pro Ala Gly Glu Arg Gly Glu Gln Gly Pro Ala Gly Ser Pro
625             630             635             640
Gly Phe Gln Gly Leu Pro Gly Pro Ala Gly Pro Pro Gly Glu Ala Gly
            645             650             655
Lys Pro Gly Glu Gln Gly Val Pro Gly Asp Leu Gly Ala Pro Gly Pro
        660             665             670
Ser Gly Ala Arg Gly Glu Arg Gly Phe Pro Gly Glu Arg Gly Val Gln
        675             680             685
Gly Pro Pro Gly Pro Ala Gly Pro Arg Gly Ala Asn Gly Ala Pro Gly
    690             695             700
Asn Asp Gly Ala Lys Gly Asp Ala Gly Ala Pro Gly Ala Pro Gly Ser
705             710             715             720
Gln Gly Ala Pro Gly Leu Gln Gly Met Pro Gly Glu Arg Gly Ala Ala
            725             730             735
Gly Leu Pro Gly Pro Lys Gly Asp Arg Gly Asp Ala Gly Pro Lys Gly
            740             745             750
Ala Asp Gly Ser Pro Gly Lys Asp Gly Val Arg Gly Leu Thr Gly Pro
    755             760             765
Ile Gly Pro Pro Gly Pro Ala Gly Ala Pro Gly Asp Lys Gly Glu Ser
    770             775             780
Gly Pro Ser Gly Pro Ala Gly Pro Thr Gly Ala Arg Gly Ala Pro Gly
785             790             795             800
Asp Arg Gly Glu Pro Gly Pro Pro Gly Pro Ala Gly Phe Ala Gly Pro
            805             810             815
Pro Gly Ala Asp Gly Gln Pro Gly Ala Lys Gly Glu Pro Gly Asp Ala
        820             825             830
Gly Ala Lys Gly Asp Ala Gly Pro Pro Gly Pro Ala Gly Pro Ala Gly
    835             840             845
Pro Pro Gly Pro Ile Gly Asn Val Gly Ala Pro Gly Ala Lys Gly Ala
```

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           850                    855                    860
Arg Gly Ser Ala Gly Pro Pro Gly Ala Thr Gly Phe Pro Gly Ala Ala
865                    870                    875                    880
Gly Arg Val Gly Pro Pro Gly Pro Ser Gly Asn Ala Gly Pro Pro Gly
                       885              890              895
Pro Pro Gly Pro Ala Gly Lys Glu Gly Gly Lys Gly Pro Arg Gly Glu
              900              905              910
Thr Gly Pro Ala Gly Arg Pro Gly Glu Val Gly Pro Pro Gly Pro Pro
         915              920              925
Gly Pro Ala Gly Glu Lys Gly Ser Pro Gly Ala Asp Gly Pro Ala Gly
    930              935              940
Ala Pro Gly Thr Pro Gly Pro Gln Gly Ile Ala Gly Gln Arg Gly Val
945                    950              955                    960
Val Gly Leu Pro Gly Gln Arg Gly Glu Arg Gly Phe Pro Gly Leu Pro
              965              970              975
Gly Pro Ser Gly Glu Pro Gly Lys Gln Gly Pro Ser Gly Ala Ser Gly
         980              985              990
Glu Arg Gly Pro Pro Gly Pro Met Gly Pro Pro Gly Leu Ala Gly Pro
         995              1000             1005
Pro Gly Glu Ser Gly Arg Glu Gly Ala Pro Ala Ala Glu Gly Ser Pro
    1010             1015             1020
Gly Arg Asp Gly Ser Pro Gly Ala Lys Gly Asp Arg Gly Glu Thr Gly
1025             1030             1035             1040
Pro Ala Gly Pro Pro Gly Ala Pro Gly Ala Pro Gly Ala Pro Gly Pro
         1045             1050             1055
Val Gly Pro Ala Gly Lys Ser Gly Asp Arg Gly Glu Thr Gly Pro Ala
         1060             1065             1070
Gly Pro Ala Gly Pro Val Gly Pro Val Gly Ala Arg Gly Pro Ala Gly
         1075             1080             1085
Pro Gln Gly Pro Arg Gly Asp Lys Gly Glu Thr Gly Glu Gln Gly Asp
    1090             1095             1100
Arg Gly Ile Lys Gly His Arg Gly Phe Ser Gly Leu Gln Gly Pro Pro
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Gly Pro Pro Gly Ser Pro Gly Glu Gln Gly Pro Ser Gly Ala Ser Gly
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Pro Ala Gly Pro Arg Gly Pro Pro Gly Ser Ala Gly Ala Pro Gly Lys
         1140             1145             1150
Asp Gly Leu Asn Gly Leu Pro Gly Pro Ile Gly Pro Pro Gly Pro Arg
         1155             1160             1165
Gly Arg Thr Gly Asp Ala Gly Pro Val Gly Pro Pro Gly Pro Pro Gly
    1170             1175             1180
Pro Pro Gly Pro Pro Gly Pro Pro Ser Ala Gly Phe Asp Phe Ser Phe
1185             1190             1195             1200
Leu Pro Gln Pro Pro Gln Glu Lys Ala His Asp Gly Gly Arg Tyr Tyr
              1205             1210             1215
Arg Ala Asp Asp Ala Asn Val Val Arg Asp Arg Asp Leu Glu Val Asp
         1220             1225             1230
Thr Thr Leu Lys Ser Leu Ser Gln Gln Ile Glu Asn Ile Arg Ser Pro
    1235             1240             1245
Glu Gly Ser Arg Lys Asn Pro Ala Arg Thr Cys Arg Asp Leu Lys Met
    1250             1255             1260
Cys His Ser Asp Trp Lys Ser Gly Glu Tyr Trp Ile Asp Pro Asn Gln
1265             1270             1275             1280
Gly Cys Asn Leu Asp Ala Ile Lys Val Phe Cys Asn Met Glu Thr Gly
              1285             1290             1295
Glu Thr Cys Val Tyr Pro Thr Gln Pro Ser Val Ala Gln Lys Asn Trp
         1300             1305             1310
Tyr Ile Ser Lys Asn Pro Lys Asp Lys Arg His Val Trp Phe Gly Glu
         1315             1320             1325
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Ser Met Thr Asp Gly Phe Gln Phe Glu Tyr Gly Gly Gln Gly Ser Asp
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Pro Ala Asp Val Ala Ile Gln Leu Thr Phe Leu Arg Leu Met Ser Thr
1345              1350              1355              1360
Glu Ala Ser Gln Asn Ile Thr Tyr His Cys Lys Asn Ser Val Ala Tyr
              1365              1370              1375
Met Asp Gln Gln Thr Gly Asn Leu Lys Lys Ala Leu Leu Leu Lys Gly
          1380              1385              1390
Ser Asn Glu Ile Glu Ile Arg Ala Glu Gly Asn Ser Arg Phe Thr Tyr
          1395              1400              1405
Ser Val Thr Val Asp Gly Cys Thr Ser His Thr Gly Ala Trp Gly Lys
    1410              1415              1420
Thr Val Ile Glu Tyr Lys Thr Thr Lys Ser Ser Arg Leu Pro Ile Ile
1425              1430              1435              1440
Asp Val Ala Pro Leu Asp Val Gly Ala Pro Asp Gln Glu Phe Gly Phe
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Asp Val Gly Pro Val Cys Phe Leu
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<211> 5086
<212> DNA
<213> Homo sapiens

<220>
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<222> 27, 46
<223> n = A,T,C or G

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gcatgtctaa gtgctagaca tgctcagctt tgtggatacg cggactttgt tgctgcttgc 180
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<210> 38

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<212> PRT
<213> Homo sapiens

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            20                  25                  30
Gly Pro Ala Gly Asp Arg Gly Pro Arg Gly Glu Arg Gly Pro Pro Gly
        35                  40                  45
Pro Pro Gly Arg Asp Gly Glu Asp Gly Pro Thr Gly Pro Pro Gly Pro
    50                  55                  60
Pro Gly Pro Pro Gly Pro Pro Gly Leu Gly Gly Asn Phe Ala Ala Gln
65                  70                  75                  80
Tyr Asp Gly Lys Gly Val Gly Leu Gly Pro Gly Pro Met Gly Leu Met
                85                  90                  95
Gly Pro Arg Gly Pro Pro Gly Ala Ala Gly Ala Pro Gly Pro Gln Gly
            100                 105                 110
Phe Gln Gly Pro Ala Gly Glu Pro Gly Glu Pro Gly Gln Thr Gly Pro
        115                 120                 125
Ala Gly Ala Arg Gly Pro Ala Gly Pro Pro Gly Lys Ala Gly Glu Asp
    130                 135                 140
Gly His Pro Gly Lys Pro Gly Arg Pro Gly Glu Arg Gly Val Val Gly
145                 150                 155                 160
Pro Gln Gly Ala Arg Gly Phe Pro Gly Thr Pro Gly Leu Pro Gly Phe
                165                 170                 175
Lys Gly Ile Arg Gly His Asn Gly Leu Asp Gly Leu Lys Gly Gln Pro
            180                 185                 190
Gly Ala Pro Gly Val Lys Gly Glu Pro Gly Ala Pro Gly Glu Asn Gly
        195                 200                 205
Thr Pro Gly Gln Thr Gly Ala Arg Gly Leu Pro Gly Glu Arg Gly Arg
    210                 215                 220
Val Gly Ala Pro Gly Pro Ala Gly Ala Arg Gly Ser Asp Gly Ser Val
225                 230                 235                 240
Gly Pro Val Gly Pro Ala Gly Pro Ile Gly Ser Ala Gly Pro Pro Gly
                245                 250                 255
Phe Pro Gly Ala Pro Gly Pro Lys Gly Glu Ile Gly Ala Val Gly Asn
            260                 265                 270
Ala Gly Pro Ala Gly Pro Ala Gly Pro Arg Gly Glu Val Gly Leu Pro
        275                 280                 285
Gly Leu Ser Gly Pro Val Gly Pro Pro Gly Asn Pro Gly Ala Asn Gly
    290                 295                 300
Leu Thr Gly Ala Lys Gly Ala Ala Gly Leu Pro Gly Val Ala Gly Ala
305                 310                 315                 320
Pro Gly Leu Pro Gly Pro Arg Gly Ile Pro Gly Pro Val Gly Ala Ala
                325                 330                 335
Gly Ala Thr Gly Ala Arg Gly Leu Val Gly Glu Pro Gly Pro Ala Gly
        340                 345                 350
Ser Lys Gly Glu Ser Gly Asn Lys Gly Glu Pro Gly Ser Ala Gly Pro
    355                 360                 365
Gln Gly Pro Pro Gly Pro Ser Gly Glu Glu Gly Lys Arg Gly Pro Asn
    370                 375                 380
Gly Glu Ala Gly Ser Ala Gly Pro Pro Gly Pro Pro Gly Leu Arg Gly
385                 390                 395                 400
Ser Pro Gly Ser Arg Gly Leu Pro Gly Ala Asp Gly Arg Ala Gly Val
                405                 410                 415
Met Gly Pro Pro Gly Ser Arg Gly Ala Ser Gly Pro Ala Gly Val Arg
            420                 425                 430
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```
Gly Pro Asn Gly Asp Ala Gly Arg Pro Gly Glu Pro Gly Leu Met Gly
        435           ·         440           445
Pro Arg Gly Leu Pro Gly Ser Pro Gly Asn Ile Gly Pro Ala Gly Lys
    450               455           460
Glu Gly Pro Val Gly Leu Pro Gly Ile Asp Gly Arg Pro Gly Pro Ile
465               470           475               480
Gly Pro Ala Gly Ala Arg Gly Glu Pro Gly Asn Ile Gly Phe Pro Gly
            485           ·    490               495
Pro Lys Gly Pro Thr Gly Asp Pro Gly Lys Asn Gly Asp Lys Gly His
        500               505           510
Ala Gly Leu Ala Gly Ala Arg Gly Ala Pro Gly Pro Asp Gly Asn Asn
    515           .       520           525
Gly Ala Gln Gly Pro Pro Gly Pro Gln Gly Val Gln Gly Gly Lys Gly
    530               535           540
Glu Gln Gly Pro Ala Gly Pro Pro Gly Phe Gln Gly Leu Pro Gly Pro
545               550           555               560
Ser Gly Pro Ala Gly Glu Val Gly Lys Pro Gly Glu Arg Gly Leu His
            565           570           575
Gly Glu Phe Gly Leu Pro Gly Pro Ala Gly Pro Arg Gly Glu Arg Gly
            580           585           590
Pro Pro Gly Glu Ser Gly Ala Ala Gly Pro Thr Gly Pro Ile Gly Ser
        595           600           605
Arg Gly Pro Ser Gly Pro Pro Gly Pro Asp Gly Asn Lys Gly Glu Pro
        610           615           620
Gly Val Val Gly Ala Val Gly Thr Ala Gly Pro Ser Gly Pro Ser Gly
625               630           635               640
Leu Pro Gly Glu Arg Gly Ala Ala Gly Ile Pro Gly Gly Lys Gly Glu
            645           650           655
Lys Gly Glu Pro Gly Leu Arg Gly Glu Ile Gly Asn Pro Gly Arg Asp
            660           665           670
Gly Ala Arg Gly Ala His Gly Ala Val Gly Ala Pro Gly Pro Ala Gly
        675           680           685
Ala Thr Gly Asp Arg Gly Glu Ala Gly Ala Ala Gly Pro Ala Gly Pro
    690               695           700
Ala Gly Pro Arg Gly Ser Pro Gly Glu Arg Gly Glu Val Gly Pro Ala
705               710           715               720
Gly Pro Asn Gly Phe Ala Gly Pro Ala Gly Ala Ala Gly Gln Pro Gly
            725           730           ·    735
Ala Lys Gly Glu Arg Gly Ala Lys Gly Pro Lys Gly Glu Asn Gly Val
            740           745           750
Val Gly Pro Thr Gly Pro Val Gly Ala Ala Gly Pro Ala Gly Pro Asn
        755           760           765
Gly Pro Pro Gly Pro Ala Gly Ser Arg Gly Asp Gly Gly Pro Pro Gly
    770               775           780
Met Thr Gly Phe Pro Gly Ala Ala Gly Arg Thr Gly Pro Pro Gly Pro
785               790           795               800
Ser Gly Ile Ser Gly Pro Pro Gly Pro Pro Gly Pro Ala Gly Lys Glu
            805           810           815
Gly Leu Arg Gly Pro Arg Gly Asp Gln Gly Pro Val Gly Arg Thr Gly
        820           825           830
Glu Val Gly Ala Val Gly Pro Pro Gly Phe Ala Gly Glu Lys Gly Pro
        835           840           845 ·
Ser Gly Glu Ala Gly Thr Ala Gly Pro Pro Gly Thr Pro Gly Pro Gln
    850 ·           855           860
Gly Leu Leu Gly Ala Pro Gly Ile Leu Gly Leu Pro Gly Ser Arg Gly
865               870           875               880
Glu Arg Gly Leu Pro Gly Val Ala Gly Ala Val Gly Glu Pro Gly Pro
            885           890           895
Leu Gly Ile Ala Gly Pro Pro Gly Ala Arg Gly Pro Pro Gly Ala Val
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           915                920                925
Asn Pro Gly Asn Asp Gly Pro Pro Gly Arg Asp Gly Gln Pro Gly His
     930                935                940
Lys Gly Glu Arg Gly Tyr Pro Gly Asn Ile Gly Pro Val Gly Ala Ala
945                950                955                960
Gly Ala Pro Gly Pro His Gly Pro Val Gly Pro Ala Gly Lys His Gly
               965                970                975
Asn Arg Gly Glu Thr Gly Pro Ser Gly Pro Val Gly Pro Ala Gly Ala
           980                985                990
Val Gly Pro Arg Gly Pro Ser Gly Pro Gln Gly Ile Arg Gly Asp Lys
           995                1000               1005
Gly Glu Pro Gly Glu Lys Gly Pro Arg Gly Leu Pro Gly Leu Lys Gly
       1010               1015               1020
His Asn Gly Leu Gln Gly Leu Pro Gly Ile Ala Gly His His Gly Asp
1025               1030               1035               1040
Gln Gly Ala Pro Gly Ser Val Gly Pro Ala Gly Pro Arg Gly Pro Ala
               1045               1050               1055
Gly Pro Ser Gly Pro Ala Gly Lys Asp Gly Arg Thr Gly His Pro Gly
           1060               1065               1070
Thr Val Gly Pro Ala Gly Ile Arg Gly Pro Gln Gly His Gln Gly Pro
           1075               1080               1085
Ala Gly Pro Pro Gly Pro Pro Gly Pro Pro Gly Pro Pro Gly Val Ser
       1090               1095               1100
Gly Gly Gly Tyr Asp Phe Gly Tyr Asp Gly Asp Phe Tyr Arg Ala Asp
1105               1110               1115               1120
Gln Pro Arg Ser Ala Pro Ser Leu Arg Pro Lys Asp Tyr Glu Val Asp
               1125               1130               1135
Ala Thr Leu Lys Ser Leu Asn Asn Gln Ile Glu Thr Leu Leu Thr Pro
           1140               1145               1150
Glu Gly Ser Arg Lys Asn Pro Ala Arg Thr Cys Arg Asp Leu Arg Leu
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Ser His Pro Glu Trp Ser Ser Gly Tyr Tyr Trp Ile Asp Pro Asn Gln
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Gly Cys Thr Met Asp Ala Ile Lys Val Tyr Cys Asp Phe Ser Thr Gly
1185               1190               1195               1200
Glu Thr Cys Ile Arg Ala Gln Pro Glu Asn Ile Pro Ala Lys Asn Trp
               1205               1210               1215
Tyr Arg Ser Ser Lys Asp Lys Lys His Val Trp Leu Gly Glu Thr Ile
               1220               1225               1230
Asn Ala Gly Ser Gln Phe Glu Tyr Asn Val Glu Gly Val Thr Ser Lys
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Glu Met Ala Thr Gln Leu Ala Phe Met Arg Leu Leu Ala Asn Tyr Ala
       1250               1255               1260
Ser Gln Asn Ile Thr Tyr His Cys Lys Asn Ser Ile Ala Tyr Met Asp
1265                1270               1275               1280
Glu Glu Thr Gly Asn Leu Lys Lys Ala Val Ile Leu Gln Gly Ser Asn
               1285               1290               1295
Asp Val Glu Leu Val Ala Glu Gly Asn Ser Arg Phe Thr Tyr Thr Val
           1300               1305               1310
Leu Val Asp Gly Cys Ser Lys Lys Thr Asn Glu Trp Gly Lys Thr Ile
       1315               1320               1325
Ile Glu Tyr Lys Thr Asn Lys Pro Ser Arg Leu Pro Phe Leu Asp Ile
       1330               1335               1340
Ala Pro Leu Asp Ile Gly Gly Ala Asp His Glu Phe Phe Val Asp Ile
1345               1350               1355               1360
Gly Pro Val Cys Phe Lys
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<210> 39
<211> 2235
<212> DNA
<213> Homo sapiens

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cctcctcaga tgccaccaca aattccacaa·taccagcccc tgggtcagca agtacctcac 180
atgcctttgg ccaaagatgg cctcgccatg ggcaaggaga tgccccactt gcagtatggc 240
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aaggaagccg ttcccaagaa aggcaaagaa ataccattag ccagtttacg aggggaacaa 360
ggtccccgtg gagagcctgg cccaagagga ccacctgggc ccctggtttt accaggtcat 420
gggatacctg gaattaaagg aaaaccaggg ccacagggat atccaggagt tggaaagcca 480
ggtatgcctg gaatgccagg gaagccagga gccatgggca tgcctggggc aaaaggagaa 540
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<210> 40
<211> 744
<212> PRT
<213> Homo sapiens

<400> 40
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Ile Ser Leu Ser Ser Ile Arg Leu Ile Gln Ala Gly Ala Tyr Tyr Gly
            20   25   30
Ile Lys Pro Leu Pro Pro Gln Ile Pro Pro Gln Met Pro Pro Gln Ile
        35   40   45

Pro Gln Tyr Gln Pro Leu Gly Gln Gln Val Pro His Met Pro Leu Ala
    50   55   60
Lys Asp Gly Leu Ala Met Gly Lys Glu Met Pro His Leu Gln Tyr Gly
65   70   75   80
Lys Glu Tyr Pro His Leu Pro Gln Tyr Met Lys Glu Ile Gln Pro Ala
                85   90   95
Pro Arg Met Gly Lys Glu Ala Val Pro Lys Lys Gly Lys Glu Ile Pro
            100   105   110
Leu Ala Ser Leu Arg Gly Glu Gln Gly Pro Arg Gly Glu Pro Gly Pro
        115   120   125
Arg Gly Pro Pro Gly Pro Pro Gly Leu Pro Gly His Gly Ile Pro Gly
    130   135   140
Ile Lys Gly Lys Pro Gly Pro Gln Gly Tyr Pro Gly Val Gly Lys Pro
145   150   155   160
Gly Met Pro Gly Met Pro Gly Lys Pro Gly Ala Met Gly Met Pro Gly
            165   170   175
Ala Lys Gly Glu Ile Gly Gln Lys Gly Glu Ile Gly Pro Met Gly Ile
            180   185   190
Pro Gly Pro Gln Gly Pro Pro Gly Pro His Gly Leu Pro Gly Ile Gly
        195   200   205
Lys Pro Gly Gly Pro Gly Leu Pro Gly Gln Pro Gly Pro Lys Gly Asp
    210   215   220
Arg Gly Pro Lys Gly Leu Pro Gly Pro Gln Gly Leu Arg Gly Pro Lys
225   230   235   240
Gly Asp Lys Gly Phe Gly Met Pro Gly Ala Pro Gly Val Lys Gly Pro
            245   250   255
Pro Gly Met His Gly Leu Pro Gly Pro Val Gly Leu Pro Gly Val Gly
            260   265   270
Lys Pro Gly Val Thr Gly Phe Pro Gly Pro Gln Gly Pro Leu Gly Lys
    275   280   285
Pro Gly Ala Pro Gly Glu Pro Gly Arg Gln Gly Pro Ile Gly Val Pro
    290   295   300
Gly Val Gln Gly Pro Pro Gly Ile Pro Gly Ile Gly Lys Pro Gly Gln
305   310   315   320
Asp Gly Ile Pro Gly Gln Pro Gly Phe Pro Gly Gly Lys Gly Glu Gln
            325   330   335
Gly Leu Pro Gly Leu Pro Gly Ala Pro Gly Leu Pro Gly Ile Gly Lys
        340   345   350
Pro Gly Phe Pro Gly Pro Lys Gly Asp Arg Gly Met Gly Gly Val Pro
        355   360   365
Gly Ala Leu Gly Pro Arg Gly Glu Lys Gly Pro Ile Gly Ser Pro Gly
    370   375   380
Ile Gly Gly Ser Pro Gly Glu Pro Gly Leu Pro Gly Ile Pro Gly Pro
385   390   395   400
Met Gly Pro Pro Gly Ala Ile Gly Phe Pro Gly Pro Lys Gly Glu Gly
            405   410   415
Gly Ile Val Gly Pro Gln Gly Pro Pro Gly Pro Lys Gly Glu Pro Gly
            420   425   430
Leu Gln Gly Phe Pro Gly Lys Pro Gly Phe Leu Gly Glu Val Gly Pro
        435   440   445
Pro Gly Met Arg Gly Phe Pro Gly Pro Ile Gly Pro Lys Gly Glu His
    450   455   460
Gly Gln Lys Gly Val Pro Gly Leu Pro Gly Val Pro Gly Leu Leu Gly
465   470   475   480
Pro Lys Gly Glu Pro Gly Ile Pro Gly Asp Gln Gly Leu Gln Gly Pro
            485   490   495
Pro Gly Ile Pro Gly Ile Gly Gly Pro Ser Gly Pro Ile Gly Pro Pro
            500   505   510
Gly Ile Pro Gly Pro Lys Gly Glu Pro Gly Leu Pro Gly Pro Pro Gly

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          515                      520                      525
Phe Pro Gly Ile Gly Lys Pro Gly Val Ala Gly Leu His Gly Pro Pro
     530                      535                      540
Gly Lys Pro Gly Ala Leu Gly Pro Gln Gly Gln Pro Gly Leu Pro Gly
545                      550                      555                      560
Pro Pro Gly Pro Pro Gly Pro Pro Gly Pro Pro Ala Val Met Pro Pro
                    565                      570                      575
Thr Pro Pro Pro Gln Gly Glu Tyr Leu Pro Asp Met Gly Leu Gly Ile
               580                      585                      590
Asp Gly Val Lys Pro Pro His Ala Thr Gly Ala Lys Lys Gly Lys Asn
          595                      600                      605
Gly Gly Pro Ala Tyr Glu Met Pro Ala Phe Thr Ala Glu Leu Thr Ala
     610                      615                      620
Pro Phe Pro Pro Val Gly Gly Pro Val Lys Phe Asn Lys Leu Leu Tyr
625                      630                      635                      640
Asn Gly Arg Gln Asn Tyr Asn Pro Gln Thr Gly Ile Phe Thr Cys Glu
                    645                      650                      655
Val Pro Gly Val Tyr Tyr Phe Ala Tyr His Val His Cys Lys Gly Gly
               660                      665                      670
Asn Val Trp Val Ala Leu Phe Lys Asn Asn Glu Pro Val Met Tyr Thr
          675                      680                      685
Tyr Asp Glu Tyr Lys Lys Gly Phe Leu Asp Gln Ala Ser Gly Ser Ala
     690                      695                      700
Val Leu Leu Leu Arg Pro Gly Asp Arg Val Phe Leu Gln Met Pro Ser
705                      710                      715                      720
Glu Gln Ala Ala Gly Leu Tyr Ala Gly Gln Tyr Val His Ser Ser Phe
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Ser Gly Tyr Leu Leu Tyr Pro Met
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<210> 41
<211> 5064
<212> DNA
<213> Homo sapiens

<400> 41
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aactcttaat catctcccat tttccttaga catttaaatt tcaaggcagg taccctctgt 4920
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ttgttctttt caactattgt tctcatattt ctcacactaa ataaatctct atgagagctt 5040
cttgaaaaaa aaaaaaaaaa agcg                                         5064
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<210> 42
<211> 1224
<212> PRT
<213> Homo sapiens

<400> 42

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Met Leu Thr Lys Phe Glu Thr Lys Ser Ala Arg Val Lys Gly Leu Ser
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Phe His Pro Lys Arg Pro Trp Ile Leu Thr Ser Leu His Asn Gly Val
            20                  25                  30
Ile Gln Leu Trp Asp Tyr Arg Met Cys Thr Leu Ile Asp Lys Phe Asp
        35                  40                  45
Glu His Asp Gly Pro Val Arg Gly Ile Asp Phe His Lys Gln Gln Pro
    50                  55                  60
Leu Phe Val Ser Gly Gly Asp Asp Tyr Lys Ile Lys Val Trp Asn Tyr
65                  70                  75                  80
Lys Leu Arg Arg Cys Leu Phe Thr Leu Leu Gly His Leu Asp Tyr Ile
                85                  90                  95
Arg Thr Thr Phe Phe His His Glu Tyr Pro Trp Ile Leu Ser Ala Ser
            100                 105                 110
Asp Asp Gln Thr Ile Arg Val Trp Asn Trp Gln Ser Arg Thr Cys Val
            115                 120                 125
Cys Val Leu Thr Gly His Asn His Tyr Val Met Cys Ala Gln Phe His
        130                 135                 140
Pro Thr Glu Asp Leu Val Val Ser Ala Ser Leu Asp Gln Thr Val Arg
145                 150                 155                 160
Val Trp Asp Ile Ser Gly Leu Arg Lys Lys Asn Leu Ser Pro Gly Ala
                165                 170                 175
Val Glu Ser Asp Val Arg Gly Ile Thr Gly Val Asp Leu Phe Gly Thr
            180                 185                 190
Thr Asp Ala Val Val Lys His Val Leu Glu Gly His Asp Arg Gly Val
            195                 200                 205
Asn Trp Ala Ala Phe His Pro Thr Met Pro Leu Ile Val Ser Gly Ala
        210                 215                 220
Asp Asp Arg Gln Val Lys Ile Trp Arg Met Asn Glu Ser Lys Ala Trp
225                 230                 235                 240
Glu Val Asp Thr Cys Arg Gly His Tyr Asn Asn Val Ser Cys Ala Val
                245                 250                 255
Phe His Pro Arg Gln Glu Leu Ile Leu Ser Asn Ser Glu Asp Lys Ser
            260                 265                 270
Ile Arg Val Trp Asp Met Ser Lys Arg Thr Gly Val Gln Thr Phe Arg
            275                 280                 285
Arg Asp His Asp Arg Phe Trp Val Leu Ala Ala His Pro Asn Leu Asn
        290                 295                 300
Leu Phe Ala Ala Gly His Asp Gly Gly Met Ile Val Phe Lys Leu Glu
305                 310                 315                 320
Arg Glu Arg Pro Ala Tyr Ala Val His Gly Asn Met Leu His Tyr Val
                325                 330                 335
Lys Asp Arg Phe Leu Arg Gln Leu Asp Phe Asn Ser Ser Lys Asp Val
            340                 345                 350
Ala Val Met Gln Leu Arg Ser Gly Ser Lys Phe Pro Val Phe Asn Met
        355                 360                 365
Ser Tyr Asn Pro Ala Glu Asn Ala Val Leu Leu Cys Thr Arg Ala Ser
370                 375                 380
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Asn Leu Glu Asn Ser Thr Tyr Asp Leu Tyr Thr Ile Pro Lys Asp Ala
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Asp Ser Gln Asn Pro Asp Ala Pro Glu Gly Lys Arg Ser Ser Gly Leu
            405             410             415
Thr Ala Val Trp Val Ala Arg Asn Arg Phe Ala Val Leu Asp Arg Met
            420             425             430
His Ser Leu Leu Ile Lys Asn Leu Lys Asn Glu Ile Thr Lys Lys Val
        435             440             445
Gln Val Pro Asn Cys Asp Glu Ile Phe Tyr Ala Gly Thr Gly Asn Leu
    450             455             460
Leu Leu Arg Asp Ala Asp Ser Ile Thr Leu Phe Asp Val Gln Gln Lys
465             470             475             480
Arg Thr Leu Ala Ser Val Lys Ile Ser Lys Val Lys Tyr Val Ile Trp
            485             490             495
Ser Ala Asp Met Ser His Val Ala Leu Leu Ala Lys His Ala Ile Val
            500             505             510
Ile Cys Asn Arg Lys Leu Asp Ala Leu Cys Asn Ile His Glu Asn Ile
        515             520             525
Arg Val Lys Ser Gly Ala Trp Asp Glu Ser Gly Val Phe Ile Tyr Thr
    530             535             540
Thr Ser Asn His Ile Lys Tyr Ala Val Thr Thr Gly Asp His Gly Ile
545             550             555             560
Ile Arg Thr Leu Asp Leu Pro Ile Tyr Val Thr Arg Val Lys Gly Asn
            565             570             575
Asn Val Tyr Cys Leu Asp Arg Glu Cys Arg Pro Arg Val Leu Thr Ile
        580             585             590
Asp Pro Thr Glu Phe Lys Phe Lys Leu Ala Leu Ile Asn Arg Lys Tyr
    595             600             605
Asp Glu Val Leu His Met Val Arg Asn Ala Lys Leu Val Gly Gln Ser
    610             615             620
Ile Ile Ala Tyr Leu Gln Lys Lys Gly Tyr Pro Glu Val Ala Leu His
.625             630             635             640
Phe Val Lys Asp Glu Lys Thr Arg Phe Ser Leu Ala Leu Glu Cys Gly
            645             650             655
Asn Ile Glu Ile Ala Leu Glu Ala Ala Lys Ala Leu Asp Asp Lys Asn
            660             665             670
Cys Trp Glu Lys Leu Gly Glu Val Ala Leu Leu Gln Gly Asn His Gln
        675             680             685
Ile Val Glu Met Cys Tyr Gln Arg Thr Lys Asn Phe Asp Lys Val Ser
    690             695             700
Phe Leu Tyr Leu Ile Thr Gly Asn Leu Glu Lys Leu Arg Lys Met Met
705             710             715             720
Lys Ile Ala Glu Ile Arg Lys Asp Met Ser Gly His Tyr Gln Asn Ala
            725             730             735
Leu Tyr Leu Gly Asp Val Ser Glu Arg Val Arg Ile Leu Lys Asn Cys
        740             745             750
Gly Gln Lys Ser Leu Ala Tyr Leu Thr Ala Ala Thr His Gly Leu Asp
    755             760             765
Glu Glu Ala Glu Ser Leu Lys Glu Thr Phe Asp Pro Glu Lys Glu Thr
    770             775             780
Ile Pro Asp Ile Asp Pro Asn Ala Lys Leu Leu Gln Pro Pro Ala Pro
785             790             795             800
Ile Met Pro Leu Asp Thr Asn Trp Pro Leu Leu Thr Val Ser Lys Gly
            805             810             815
Phe Phe Glu Gly Thr Ile Ala Ser Lys Gly Lys Gly Gly Ala Leu Ala
            820             825             830
Ala Asp Ile Asp Ile Asp Thr Val Gly Thr Glu Gly Trp Gly Glu Asp
        835             840             845
Ala Glu Leu Gln Leu Asp Glu Asp Gly Phe Val Glu Ala Thr Glu Gly
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Leu Gly Asp Asp Ala Leu Gly Lys Gly Gln Glu Glu Gly Gly Gly Trp
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Asp Val Glu Glu Asp Leu Glu Leu Pro Pro Glu Leu Asp Ile Ser Pro
                  885                   890                   895
Gly Ala Ala Gly Gly Ala Glu Asp Gly Phe Phe Val Pro Pro Thr Lys
                  900                   905                   910
Gly Thr Ser Pro Thr Gln Ile Trp Cys Asn Asn Ser Gln Leu Pro Val
              915                   920                   925
Asp His Ile Leu Ala Gly Ser Phe Glu Thr Ala Met Arg Leu Leu His
          930                   935                   940
Asp Gln Val Gly Val Ile Gln Phe Gly Pro Tyr Lys Gln Leu Phe Leu
945                   950                   955                   960
Gln Thr Tyr Ala Arg Gly Arg Thr Thr Tyr Gln Ala Leu Pro Cys Leu
                  965                   970                   975
Pro Ser Met Tyr Gly Tyr Pro Asn Arg Asn Trp Lys Asp Ala Gly Leu
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Lys Asn Gly Val Pro Ala Val Gly Leu Lys Leu Asn Asp Leu Ile Gln
              995                   1000                  1005
Arg Leu Gln Leu Cys Tyr Gln Leu Thr Thr Val Gly Lys Phe Glu Glu
          1010                  1015                  1020
Ala Val Glu Lys Phe Arg Ser Ile Leu Leu Ser Val Pro Leu Leu Val
1025                  1030                  1035                  1040
Val Asp Asn Lys Gln Glu Ile Ala Glu Ala Gln Gln Leu Ile Thr Ile
                  1045                  1050                  1055
Cys Arg Glu Tyr Ile Val Gly Leu Ser Val Glu Thr Glu Arg Lys Lys
              1060                  1065                  1070
Leu Pro Lys Glu Thr Leu Glu Gln Gln Lys Arg Ile Cys Glu Met Ala
              1075                  1080                  1085
Ala Tyr Phe Thr His Ser Asn Leu Gln Pro Val His Met Ile Leu Val
          1090                  1095                  1100
Leu Arg Thr Ala Leu Asn Leu Phe Phe Lys Leu Lys Asn Phe Lys Thr
1105                  1110                  1115                  1120
Ala Ala Thr Phe Ala Arg Arg Leu Leu Glu Leu Gly Pro Lys Pro Glu
                  1125                  1130                  1135
Val Ala Gln Gln Thr Arg Lys Ile Leu Ser Ala Cys Glu Lys Asn Pro
              1140                  1145                  1150
Thr Asp Ala Tyr Gln Leu Asn Tyr Asp Met His Asn Pro Phe Asp Ile
          1155                  1160                  1165
Cys Ala Ala Ser Tyr Arg Pro Ile Tyr Arg Gly Lys Pro Val Glu Lys
          1170                  1175                  1180
Cys Pro Leu Ser Gly Ala Cys Tyr Ser Pro Glu Phe Lys Gly Gln Ile
1185                  1190                  1195                  1200
Cys Arg Val Thr Thr Val Thr Glu Ile Gly Lys Asp Val Ile Gly Leu
                  1205                  1210                  1215
Arg Ile Ser Pro Leu Gln Phe Arg
                  1220
```

<210> 43
<211> 266
<212> DNA
<213> Homo sapiens

<400> 43
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atgcccaagt gtcccaagtg caacaaggag gtgtacttcg ccgagagggt gacctctctg 60
ggcaaggact ggcatcggcc ctgcctgaag tgcgagaaat gtgggaagac gctgacctct 120
gggggccacg ctgagcacga aggcaaaccc tactgcaacc accctgctca cgcagccatg 180
tttgggccta aaggctttgg gcggggcgga gccgagagcc acactttcaa gtaaaccagg 240
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tggtggagac ccatccttgg ctgctt   266

<210> 44
<211> 77
<212> PRT
<213> Homo sapiens

<400> 44
Met Pro Lys Cys Pro Lys Cys Asn Lys Glu Val Tyr Phe Ala Glu Arg
 1   5   10   15
Val Thr Ser Leu Gly Lys Asp Trp His Arg Pro Cys Leu Lys Cys Glu
                20   25   30
Lys Cys Gly Lys Thr Leu Thr Ser Gly Gly His Ala Glu His Glu Gly
            35   40   45
Lys Pro Tyr Cys Asn His Pro Cys Tyr Ala Ala Met Phe Gly Pro Lys
        50   55   60
Gly Phe Gly Arg Gly Gly Ala Glu Ser His Thr Phe Lys
65   70   75

<210> 45
<211> 2312
<212> DNA
<213> Homo sapiens

<400> 45
tccagtgacg gagccgcccg gccgacagcc ccgagacgac agcccggcgc gtcccggtcc 60
ccacctccga ccaccgccag cgctccaggc cccgcgctcc ccgctcgccg ccaccgcgcc 120
ctccgctccg cccgcagtgc caaccatgac cgccgccagt atgggccccg tccgcgtcgc 180
cttcgtggtc ctcctcgccc tctgcagccg gccggccgtc ggccagaact gcagcgggcc 240
gtgccggtgc ccggacgagc cggcgccgcg ctgcccggcg ggcgtgagcc tcgtgctgga 300
cggctgcggc tgctgccgcg tctgcgccaa gcagctgggc gagctgtgca ccgagcgcga 360
cccctgcgac ccgcacaagg cctcttctg tgacttcggc tccccggcca accgcaagat 420
cggcgtgtgc accgccaaag atggtgctcc ctgcatcttc ggtggtacgg tgtaccgcag 480
cggagagtcc ttccagagca gctgcaagta ccagtgcacg tgcctggacg gggcggtggg 540
ctgcatgccc ctgtgcagca tggacgttcg tctgcccagc cctgactgcc ccttcccgag 600
gagggtcaag ctgcccggga aatgctgcga ggagtgggtg tgtgacgagc caaggacca 660
aaccgtggtt gggcctgccc tcgcggctta ccgactggaa gacacgtttg gcccagaccc 720
aactatgatt agagccaact gcctggtcca gaccacagag tggagcgcct gttccaagac 780
ctgtgggatg ggcatctcca cccggggttac caatgacaac gcctcctgca ggctagagaa 840
gcagagccgc ctgtgcatgg tcaggccttg cgaagctgac ctggaagaga acattaagaa 900
gggcaaaaag tgcatccgta ctcccaaaat ctccaagcct atcaagtttg agctttctgg 960
ctgcaccagc atgaagacat accgagctaa attctgtgga gtatgtaccg acggccgatg 1020
ctgcacccc cacagaacca ccaccctgcc ggtggagttc aagtgccctg acggcgaggt 1080
catgaagaag aacatgatgt tcatcaagac ctgtgcctgc cattacaact gtcccggaga 1140
caatgacatc tttgaatcgc tgtactacag gaagatgtac ggagacatgg catgaagcca 1200
gagagtgaga gacattaact cattagactg gaacttgaac tgattcacat ctcattttc 1260
cgtaaaaatg atttcagtag cacaagttat ttaaatctgt ttttctaact gggggaaaag 1320
attcccaccc aattcaaaac attgtgccat gtcaaacaaa tagtctatct tccccagaca 1380
ctggtttgaa gaatgttaag acttgacagt ggaactacat tagtacacag caccagaatg 1440
tatattaagg tgtggcttta ggagcagtgg gagggtacca gcagaaaggt tagtatcatc 1500
agatagctct tatacgagta atatgcctgc tatttgaagt gtaattgaga aggaaaattt 1560
tagcgtgctc actgacctgc ctgtagcccc agtgacagct aggatgtgca ttctccagcc 1620
atcaagagac tgagtcaagt tgttccttaa gtcagaacag cagactcagc tctgacattc 1680
tgattcgaat gacactgttc aggaatcgga atcctgtcga ttagactgga cagcttgtgg 1740
caagtgaatt tcctgtaaca agccagattt tttaaaattt atattgtaaa tattgtgtgt 1800
gtgtgtgtgt gtgtatatat atatatat gtacagttat ctaagttaat ttaaagttgt 1860
ttgtgccttt ttatttttgt ttttaatgct ttgatatttc aatgttagcc tcaatttctg 1920
aacaccatag gtagaatgta aagcttgtct gatcgttcaa agcatgaaat ggatacttat 1980

```
atggaaattc tctcagatag aatgacagtc cgtcaaaaca gattgtttgc aaaggggagg 2040
catcagtgtc cttggcaggc tgatttctag gtaggaaatg tggtagctca cgctcacttt 2100
taatgaacaa atggccttta ttaaaaactg agtgactcta tatagctgat cagttttttc 2160
acctggaagc atttgtttct actttgatat gactgttttt cggacagttt atttgttgag 2220
agtgtgacca aaagttacat gtttgcacct ttctagttga aaataaagta tatttttttct 2280
aaaaaaaaaa aaaaacgaca gcaacggaat tc                                2312
```

<210> 46
<211> 349
<212> PRT
<213> Homo sapiens

<400> 46

```
Met Thr Ala Ala Ser Met Gly Pro Val Arg Val Ala Phe Val Val Leu
1               5                   10                  15
Leu Ala Leu Cys Ser Arg Pro Ala Val Gly Gln Asn Cys Ser Gly Pro
                20                  25                  30
Cys Arg Cys Pro Asp Glu Pro Ala Pro Arg Cys Pro Ala Gly Val Ser
            35                  40                  45
Leu Val Leu Asp Gly Cys Gly Cys Cys Arg Val Cys Ala Lys Gln Leu
        50                  55                  60
Gly Glu Leu Cys Thr Glu Arg Asp Pro Cys Asp Pro His Lys Gly Leu
65                  70                  75                  80
Phe Cys Asp Phe Gly Ser Pro Ala Asn Arg Lys Ile Gly Val Cys Thr
                85                  90                  95
Ala Lys Asp Gly Ala Pro Cys Ile Phe Gly Gly Thr Val Tyr Arg Ser
                100                 105                 110
Gly Glu Ser Phe Gln Ser Ser Cys Lys Tyr Gln Cys Thr Cys Leu Asp
            115                 120                 125
Gly Ala Val Gly Cys Met Pro Leu Cys Ser Met Asp Val Arg Leu Pro
        130                 135                 140
Ser Pro Asp Cys Pro Phe Pro Arg Arg Val Lys Leu Pro Gly Lys Cys
145                 150                 155                 160
Cys Glu Glu Trp Val Cys Asp Glu Pro Lys Asp Gln Thr Val Val Gly
                165                 170                 175
Pro Ala Leu Ala Ala Tyr Arg Leu Glu Asp Thr Phe Gly Pro Asp Pro
                180                 185                 190
Thr Met Ile Arg Ala Asn Cys Leu Val Gln Thr Thr Glu Trp Ser Ala
            195                 200                 205
Cys Ser Lys Thr Cys Gly Met Gly Ile Ser Thr Arg Val Thr Asn Asp
        210                 215                 220
Asn Ala Ser Cys Arg Leu Glu Lys Gln Ser Arg Leu Cys Met Val Arg
225                 230                 235                 240
Pro Cys Glu Ala Asp Leu Glu Glu Asn Ile Lys Lys Gly Lys Lys Cys
                245                 250                 255
Ile Arg Thr Pro Lys Ile Ser Lys Pro Ile Lys Phe Glu Leu Ser Gly
                260                 265                 270
Cys Thr Ser Met Lys Thr Tyr Arg Ala Lys Phe Cys Gly Val Cys Thr
            275                 280                 285
Asp Gly Arg Cys Cys Thr Pro His Arg Thr Thr Thr Leu Pro Val Glu
        290                 295                 300
Phe Lys Cys Pro Asp Gly Glu Val Met Lys Lys Asn Met Met Phe Ile
305                 310                 315                 320
Lys Thr Cys Ala Cys His Tyr Asn Cys Pro Gly Asp Asn Asp Ile Phe
                325                 330                 335
Glu Ser Leu Tyr Tyr Arg Lys Met Tyr Gly Asp Met Ala
                340                 345
```

```
<210> 47
<211> 3025
<212> DNA
<213> Homo sapiens

<400> 47
gcacgagcag gcagttcaga ttaaagaagc taattgatca agaaatcaag tctcaggagg 60
agaaggagca agaaaaggag aaaagggtca ccaccctgaa agaggagctg accaagctga 120
agtcttttgc tttgatggtg gtggatgaac agcaaaggct gacggcacag ctcacccttc 180
aaagacagaa aatccaagag ctgaccacaa atgcaaagga aacacatacc aaactagccc 240
ttgctgaagc cagagttcag gaggaagagc agaaggcaac cagactagag aaggaactgc 300
aaacgcagac cacaaagttt caccaagacc aagacacaat tatggcgaag ctcaccaatg 360
aggacagtca aaatcgccag cttcaacaaa agctggcagc actcagccgg cagattgatg 420
agttagaaga gacaaacagg tctttacgaa aagcagaaga ggagctgcaa gatataaaag 480
aaaaaatcag taagggagaa tatggaaacg ctggtatcat ggctgaagtg gaagagctca 540
taaaaatgga ggagcagtgc agagatctca ataagaggct tgaaagggag acgttacaga 600
gtaaagactt taaactagag gttgaaaaac tcagtaaaag aattatggct ctggaaaagt 660
tagaagacgc tttcaacaaa agcaaacaag aatgctactc tctgaaatgc aatttagaaa 720
aagaaaggat gaccacaaag cagttgtctc aagaactgga gagtttaaaa gtaaggatca 780
aagagctaga agccattgaa agtcggctag aaaagacaga attcactcta aaagaggatt 840
taactaaact gaaaacatta actgtgatgt ttgtagatga acggaaaaca atgagtgaaa 900
aattaaagaa aactgaagat aaattacaag ctgcttcttc tcagcttcaa gtggagcaaa 960
ataaagtaac aacagttact gagaagttaa ttgaggaaac taaaagggcg ctcaagtcca 1020
aaaccgatgt agaagaaaag atgtacagcg taaccaagga gagagatgat ttaaaaaaca 1080
aattgaaagc ggaagaagag aaaggaaatg atctcctgtc aagagttaat atgttgaaaa 1140
ataggcttca atcattggaa gcaattgaga aagatttcct aaaaaacaaa ttaaatcaag 1200
actctgggaa atccacaaca gcattacacc aagaaaacaa taagattaag gagctctctc 1260
aagaagtgga aagactgaaa ctgaagctaa aggacatgaa agccattgag gatgacctca 1320
tgaaaacaga agatgaatat gagactctag aacgaaggta tgctaatgaa cgagacaaag 1380
ctcaattttt atctaaagag ctagaacatg ttaaaatgga acttgctaag tacaagttag 1440
cagaaaagac agagaccagc catgaacaat ggcttttcaa aaggcttcaa gaagaagaag 1500
ctaagtcagg gcacctctca agagaagtgg atgcattaaa agagaaaatt catgaataca 1560
tggcaactga agacctaata tgtcacctcc agggagatca ctcagtctgc aaaaaaaaac 1620
taaatcaaca agaaaacagg aacagagatt taggaagaga gattgaaaac ctcactaagg 1680
agttagagag gtaccggcat ttcagtaaga gcctcaggcc tagtctcaat ggaagaagaa 1740
tttccgatcc tcaagtattt tctaaagaag ttcagacaga agcagtagac aatgaaccac 1800
ctgattacaa gagcctcatt cctctggaac gtgcagtcat caatggtcag ttatatgagg 1860
agagtgagaa tcaagacgag gaccctaatg atgagggatc tgtgctgtcc ttcaaatgca 1920
gccagtctac tccatgtcct gttaacagaa agctatggat tccctggatg aaatccaagg 1980
agggccatct tcagaatgga aaaatgcaaa ctaaacccaa tgccaacttt gtgcaacctg 2040
gagatctagt cctaagccac acacctgggc agccacttca tataaaggtt actccagacc 2100
atgtacaaaa cacagccact cttgaaatca caagtccaac cacagagagt cctcactctt 2160
acacgagtac tgcagtgata ccgaactgtg gcacgccaaa gcaaaggata accatcctcc 2220
aaaacgcctc cataacacca gtaaagtcca aaacctctac cgaagacctc atgaatttag 2280
aacaaggcat gtccccaatt accatggcaa cctttgccag agcacagacc ccagagtctt 2340
gtggttctct aactccagaa aggacaatgt ccctattcag gtttttggctg tgactggttc 2400
agctagctct cctgagcagg gacgctcccc agaaccaaca gaaatcagtg ccaagcatgc 2460
gatattcaga gtctccccag accggcagtc atcatggcag tttcagcgtt caaacagcaa 2520
tagctcaagt gtgataacta ctgaggataa taaaatccac attcacttag gaagtcctta 2580
catgcaagct gtagccagcc cttcagcacc actgcaggat aaccgaactc aaggcttaat 2640
taacgggca ctaaacaaaa caaccaataa agtcaccagc agtattacta tcacaccaac 2700
agccacacct cttcctcgac aatcacaaat tacagtaagt aatatatata actgaccacg 2760
ctcaccctca tccagtccat actgatattt ttgcaaggaa ctcaatcctt ttttaatcat 2820
ccctccatat cccccaagac tgactgaact cgtactttgg gaaggtttgt gcatgaacta 2880
tacaagagta tctgaaacta actgttgcct gcatagtcat atcgagtgtg cacttactgt 2940
atatctttc atttacatac ttgtatggaa aatatttagt ctgcacttgt ataaatacat 3000
ctttatgtat ttgaaaaaaa aaaaa 3025

<210> 48
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<211> 752
<212> PRT
<213> Homo sapiens

<400> 48
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Met Val Val Asp Glu Gln Gln Arg Leu Thr Ala Gln Leu Thr Leu Gln
 1               5                  10                  15
Arg Gln Lys Ile Gln Glu Leu Thr Thr Asn Ala Lys Glu Thr His Thr
             20                  25                  30
Lys Leu Ala Leu Ala Glu Ala Arg Val Gln Glu Glu Glu Gln Lys Ala
         35                  40                  45
Thr Arg Leu Glu Lys Glu Leu Gln Thr Gln Thr Thr Lys Phe His Gln
     50                  55                  60
Asp Gln Asp Thr Ile Met Ala Lys Leu Thr Asn Glu Asp Ser Gln Asn
65                  70                  75                  80
Arg Gln Leu Gln Gln Lys Leu Ala Ala Leu Ser Arg Gln Ile Asp Glu
                 85                  90                  95
Leu Glu Glu Thr Asn Arg Ser Leu Arg Lys Ala Glu Glu Glu Leu Gln
                100                 105                 110
Asp Ile Lys Glu Lys Ile Ser Lys Gly Glu Tyr Gly Asn Ala Gly Ile
            115                 120                 125
Met Ala Glu Val Glu Glu Leu Ile Lys Met Glu Glu Gln Cys Arg Asp
    130                 135                 140
Leu Asn Lys Arg Leu Glu Arg Glu Thr Leu Gln Ser Lys Asp Phe Lys
145                 150                 155                 160
Leu Glu Val Glu Lys Leu Ser Lys Arg Ile Met Ala Leu Glu Lys Leu
                165                 170                 175
Glu Asp Ala Phe Asn Lys Ser Lys Gln Glu Cys Tyr Ser Leu Lys Cys
            180                 185                 190
Asn Leu Glu Lys Glu Arg Met Thr Thr Lys Gln Leu Ser Gln Glu Leu
            195                 200                 205
Glu Ser Leu Lys Val Arg Ile Lys Glu Leu Glu Ala Ile Glu Ser Arg
    210                 215                 220
Leu Glu Lys Thr Glu Phe Thr Leu Lys Glu Asp Leu Thr Lys Leu Lys
225                 230                 235                 240
Thr Leu Thr Val Met Phe Val Asp Glu Arg Lys Thr Met Ser Glu Lys
                245                 250                 255
Leu Lys Lys Thr Glu Asp Lys Leu Gln Ala Ala Ser Ser Gln Leu Gln
            260                 265                 270
Val Glu Gln Asn Lys Val Thr Thr Val Thr Glu Lys Leu Ile Glu Glu
    275                 280                 285
Thr Lys Arg Ala Leu Lys Ser Lys Thr Asp Val Glu Glu Lys Met Tyr
    290                 295                 300
Ser Val Thr Lys Glu Arg Asp Asp Leu Lys Asn Lys Leu Lys Ala Glu
305                 310                 315                 320
Glu Glu Lys Gly Asn Asp Leu Leu Ser Arg Val Asn Met Leu Lys Asn
            325                 330                 335
Arg Leu Gln Ser Leu Glu Ala Ile Glu Lys Asp Phe Leu Lys Asn Lys
            340                 345                 350
Leu Asn Gln Asp Ser Gly Lys Ser Thr Thr Ala Leu His Gln Glu Asn
    355                 360                 365
Asn Lys Ile Lys Glu Leu Ser Gln Glu Val Glu Arg Leu Lys Leu Lys
    370                 375                 380
Leu Lys Asp Met Lys Ala Ile Glu Asp Asp Leu Met Lys Thr Glu Asp
385                 390                 395                 400
Glu Tyr Glu Thr Leu Glu Arg Arg Tyr Ala Asn Glu Arg Asp Lys Ala
                405                 410                 415
Gln Phe Leu Ser Lys Glu Leu Glu His Val Lys Met Glu Leu Ala Lys
            420                 425                 430
```

```
Tyr Lys Leu Ala Glu Lys Thr Glu Thr Ser His Glu Gln Trp Leu Phe
        435                 440                 445
Lys Arg Leu Gln Glu Glu Glu Ala Lys Ser Gly His Leu Ser Arg Glu
        450                 455                 460
Val Asp Ala Leu Lys Glu Lys Ile His Glu Tyr Met Ala Thr Glu Asp
465                 470                 475                 480
Leu Ile Cys His Leu Gln Gly Asp His Ser Val Cys Lys Lys Lys Leu
                485                 490                 495
Asn Gln Gln Glu Asn Arg Asn Arg Asp Leu Gly Arg Glu Ile Glu Asn
            500                 505                 510
Leu Thr Lys Glu Leu Glu Arg Tyr Arg His Phe Ser Lys Ser Leu Arg
            515                 520                 525
Pro Ser Leu Asn Gly Arg Arg Ile Ser Asp Pro Gln Val Phe Ser Lys
        530                 535                 540
Glu Val Gln Thr Glu Ala Val Asp Asn Glu Pro Pro Asp Tyr Lys Ser
545                 550                 555                 560
Leu Ile Pro Leu Glu Arg Ala Val Ile Asn Gly Gln Leu Tyr Glu Glu
                565                 570                 575
Ser Glu Asn Gln Asp Glu Asp Pro Asn Asp Glu Gly Ser Val Leu Ser
            580                 585                 590
Phe Lys Cys Ser Gln Ser Thr Pro Cys Pro Val Asn Arg Lys Leu Trp
        595                 600                 605
Ile Pro Trp Met Lys Ser Lys Glu Gly His Leu Gln Asn Gly Lys Met
        610                 615                 620
Gln Thr Lys Pro Asn Ala Asn Phe Val Gln Pro Gly Asp Leu Val Leu
625                 630                 635                 640
Ser His Thr Pro Gly Gln Pro Leu His Ile Lys Val Thr Pro Asp His
                645                 650                 655
Val Gln Asn Thr Ala Thr Leu Glu Ile Thr Ser Pro Thr Thr Glu Ser
            660                 665                 670
Pro His Ser Tyr Thr Ser Thr Ala Val Ile Pro Asn Cys Gly Thr Pro
        675                 680                 685
Lys Gln Arg Ile Thr Ile Leu Gln Asn Ala Ser Ile Thr Pro Val Lys
        690                 695                 700
Ser Lys Thr Ser Thr Glu Asp Leu Met Asn Leu Glu Gln Gly Met Ser
705                 710                 715                 720
Pro Ile Thr Met Ala Thr Phe Ala Arg Ala Gln Thr Pro Glu Ser Cys
                725                 730                 735
Gly Ser Leu Thr Pro Glu Arg Thr Met Ser Leu Phe Arg Phe Trp Leu
        740                 745                 750
```

<210> 49
<211> 1480
<212> DNA
<213> Homo sapiens

<400> 49
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gcggagaaag ccagtgggaa cccagaccca taggagaccc gcgtccccgc tcggcctggc 60
caggccccgc gctatggagt tcctctgggc ccctctcttg ggtctgtgct gcagtctggc 120
cgctgctgat cgccacaccg tcttctggaa cagttcaaat cccaagttcc ggaatgagga 180
ctacaccata catgtgcagc tgaatgacta cgtggacatc atctgtccgc actatgaaga 240
tcactctgtg gcagacgctg ccatggagca gtacatactg tacctggtgg agcatgagga 300
gtaccagctg tgccagcccc agtccaagga ccaagtccgc tggcagtgca accggcccag 360
tgccaagcat ggcccggaga agctgtctga gaagttccag cgcttcacac ctttcaccct 420
gggcaaggag ttcaaagaag acacagctac tactacatc tccaaaccca tccaccagca 480
tgaagaccgc tgcttgaggt tgaaggtgac tgtcagtggc aaaatcactc acagtcctca 540
ggcccatgtc aatccacagg agaagagact tgcagcagat gacccagagg tgcgggttct 600
acatagcatc ggtcacagtg ctgccccacg cctcttccca cttgcctgga ctgtgctgct 660
```

```
ccttccactt ctgctgctgc aaaccccgtg aaggtgtatg ccacacctgg ccttaaagag 720
ggacaggctg aagagaggga caggcactcc aaacctgtct tggggccact ttcagagccc 780
ccagccctgg gaaccactcc caccacaggc ataagctatc acctagcagc ctcaaaacgg 840
gtcagtatta aggttttcaa ccggaaggag gccaaccagc ccgacagtgc catccccacc 900
ttcacctcgg agggacggag aaagaagtgg agacagtcct ttcccaccat tcctgccttt 960
aagccaaaga aacaagctgt gcaggcatgg tcccttaagg cacagtggga gctgagctgg 1020
aaggggccac gtggatgggc aaagcttgtc aaagatgccc cctccaggag agagccagga 1080
tgcccagatg aactgactga aggaaaagca agaaacagtt tcttgcttgg aagccaggta 1140
caggagaggc agcatgcttg ggctgaccca gcatctccca gcaagacctc atctgtggag 1200
ctgccacaga gaagtttgta gccaggtact gcattctctc ccatcctggg gcagcactcc 1260
ccagagctgt gccagcaggg gggctgtgcc aacctgttct tagagtgtag ctgtaagggc 1320
agtgcccatg tgtacattct gcctagagtg tagcctaaag ggcagggccc acgtgtatag 1380
tatctgtata taagttgctg tgtgtctgtc ctgatttcta caactggagt tttttttatac 1440
aatgttcttt gtctcaaaat aaagcaatgt gttttttcgg                         1480
```

```
<210> 50
<211> 205
<212> PRT
<213> Homo sapiens
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<400> 50
Met Glu Phe Leu Trp Ala Pro Leu Leu Gly Leu Cys Cys Ser Leu Ala
1               5                   10                  15
Ala Ala Asp Arg His Thr Val Phe Trp Asn Ser Ser Asn Pro Lys Phe
            20                  25                  30
Arg Asn Glu Asp Tyr Thr Ile His Val Gln Leu Asn Asp Tyr Val Asp
        35                  40                  45
Ile Ile Cys Pro His Tyr Glu Asp His Ser Val Ala Asp Ala Ala Met
    50                  55                  60
Glu Gln Tyr Ile Leu Tyr Leu Val Glu His Glu Glu Tyr Gln Leu Cys
65                  70                  75                  80
Gln Pro Gln Ser Lys Asp Gln Val Arg Trp Gln Cys Asn Arg Pro Ser
                85                  90                  95
Ala Lys His Gly Pro Glu Lys Leu Ser Glu Lys Phe Gln Arg Phe Thr
            100                 105                 110
Pro Phe Thr Leu Gly Lys Glu Phe Lys Glu Gly His Ser Tyr Tyr Tyr
        115                 120                 125
Ile Ser Lys Pro Ile His Gln His Glu Asp Arg Cys Leu Arg Leu Lys
    130                 135                 140
Val Thr Val Ser Gly Lys Ile Thr His Ser Pro Gln Ala His Val Asn
145                 150                 155                 160
Pro Gln Glu Lys Arg Leu Ala Ala Asp Asp Pro Glu Val Arg Val Leu
                165                 170                 175
His Ser Ile Gly His Ser Ala Ala Pro Arg Leu Phe Pro Leu Ala Trp
            180                 185                 190
Thr Val Leu Leu Leu Pro Leu Leu Leu Leu Gln Thr Pro
        195                 200                 205
```

```
<210> 51
<211> 15952
<212> DNA
<213> Homo sapiens
```

```
<400> 51
ccagccgtgt gtgatgagtg gccacacctt gcctcctctt cccgtcccag gcaccaacag 60
cacagagcag gccagtgtac ccagagccat ggcagccacg ctgggagccg gcacgccccc 120
caggccccag gccaggagca tagctggggt gtatgtggag gcctcgggcc aggcccagag 180
tgtctacgcc gccatggagc agggcctcct gcctgctggg ctcgggcagg ctctgctaga 240
```

```
ggcccaggca gccactgggg gcctggtgga cctcgcccgg ggccagctgc tccctgtgtc 300
caaggccctg cagcagggtc tggtgggggct ggagctgaag gagaagctgc tggccgctga 360
gcgtgccact acgggctatc ctgacccta cggcggtgag aagctggccc tctttcaggc 420
catcgggaag gaggttgtgg acagggccct ggggcagagc tggctggagg tccaactggc 480
cactggggggc ctggtggacc ccgcccaggg agtgctcgtg gccctgagc cagcctgcca 540
ccagggcctc ctggaccggg agacatggca caagctgtca gagcttgagc ctggcacagg 600
tgacctgcgc ttcctcaacc ccaacacgct gggagcggctg acataccacc agctgctgga 660
aaggtgtgtg cgtgcccccg ggtcggggct agccttgctg ccctcaaga tcaccttccg 720
ctccatgggc ggggcggtga gtgcagctga gctgctggag gtgggcatcc tggacgagca 780
ggctgtgcag ggtctgcggg agggcaggct ggccgcagtg gacgtgagtg cacgtgccga 840
ggtgcggcgc tacctggagg gtaccggcag cgtggcccgg gttgtcctgc tgcccgaagg 900
ccacaagaag agcttttttcc aggctgccac cgagcaccтg ctcccaatgg gcaccgcgct 960
gccactccta gaggcccagg ctgccaccca caccctggtg gaccccatca caggccagcg 1020 ·
gctgtgggta gacgaggcag tcagggcggg cctggtcagc ccagagctcc atgagcagct 1080
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gctgtgctgg ccaaaagcat gggctttatt ctctccattg gctgctgctc caccttagag 15840
gtgtgacctc actagcgttg actgagcgag tctgttgtgg agaagaactt tttgtagtaa 15900
tttactagga aaaattctga acaagtaaaa tatgaaggaa aaaaaaaaa aa        15952
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<210> 52
<211> 5065
<212> PRT
<213> Homo sapiens

<400> 52

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Met Ala Ala Thr Leu Gly Ala Gly Thr Pro Pro Arg Pro Gln Ala Arg
1                 5                 10                15
Ser Ile Ala Gly Val Tyr Val Glu Ala Ser Gly Gln Ala Gln Ser Val
                20                25                30
Tyr Ala Ala Met Glu Gln Gly Leu Leu Pro Ala Gly Leu Gly Gln Ala
        35                40                45
Leu Leu Glu Ala Gln Ala Ala Thr Gly Gly Leu Val Asp Leu Ala Arg
    50                55                60
Gly Gln Leu Leu Pro Val Ser Lys Ala Leu Gln Gln Gly Leu Val Gly
65                70                75                80
Leu Glu Leu Lys Glu Lys Leu Leu Ala Ala Glu Arg Ala Thr Thr Gly
                85                90                95
Tyr Pro Asp Pro Tyr Gly Gly Glu Lys Leu Ala Leu Phe Gln Ala Ile
            100               105               110
Gly Lys Glu Val Val Asp Arg Ala Leu Gly Gln Ser Trp Leu Glu Val
        115               120               125
Gln Leu Ala Thr Gly Gly Leu Val Asp Pro Ala Gln Gly Val Leu Val
    130               135               140
Ala Pro Glu Pro Ala Cys His Gln Gly Leu Leu Asp Arg Glu Thr Trp
145               150               155               160
His Lys Leu Ser Glu Leu Glu Pro Gly Thr Gly Asp Leu Arg Phe Leu
            165               170               175
Asn Pro Asn Thr Leu Glu Arg Leu Thr Tyr His Gln Leu Leu Glu Arg
            180               185               190
Cys Val Arg Ala Pro Gly Ser Gly Leu Ala Leu Leu Pro Leu Lys Ile
        195               200               205
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Thr Phe Arg Ser Met Gly Gly Ala Val Ser Ala Ala Glu Leu Leu Glu
    210             215             220
Val Gly Ile Leu Asp Glu Gln Ala Val Gln Gly Leu Arg Glu Gly Arg
225             230             235             240
Leu Ala Ala Val Asp Val Ser Ala Arg Ala Glu Val Arg Arg Tyr Leu
            245             250             255
Glu Gly Thr Gly Ser Val Ala Gly Val Val Leu Leu Pro Glu Gly His
            260             265             270
Lys Lys Ser Phe Phe Gln Ala Ala Thr Glu His Leu Leu Pro Met Gly
        275             280             285
Thr Ala Leu Pro Leu Leu Glu Ala Gln Ala Ala Thr His Thr Leu Val
    290             295             300
Asp Pro Ile Thr Gly Gln Arg Leu Trp Val Asp Glu Ala Val Arg Ala
305             310             315             320
Gly Leu Val Ser Pro Glu Leu His Glu Gln Leu Leu Val Ala Glu Gln
            325             330             335
Ala Val Thr Gly His His Asp Pro Phe Ser Gly Ser Gln Ile Pro Leu
            340             345             350
Phe Gln Ala Met Lys Lys Gly Leu Val Asp Arg Pro Leu Ala Leu Arg
        355             360             365
Leu Leu Asp Ala Gln Leu Ala Thr Gly Gly Leu Val Cys Pro Ala Arg
    370             375             380
Arg Leu Arg Leu Pro Leu Glu Ala Ala Leu Arg Cys Gly Cys Leu Asp
385             390             395             400
Glu Asp Thr Gln Arg Gln Leu Ser Gln Ala Gly Ser Phe Ser Asp Gly
            405             410             415
Thr His Gly Gly Leu Arg Tyr Glu Gln Leu Leu Ala Leu Cys Val Thr
        420             425             430
Asp Pro Glu Thr Gly Leu Ala Phe Leu Pro Leu Ser Gly Gly Pro Arg
        435             440             445
Gly Gly Glu Pro Gln Gly Pro Pro Phe Ile Lys Tyr Ser Thr Arg Gln
    450             455             460
Ala Leu Ser Thr Ala Thr Ala Thr Val Ser Val Gly Lys Phe Arg Gly
465             470             475             480
Arg Pro Val Ser Leu Trp Glu Leu Leu Phe Ser Glu Ala Ile Ser Ser
            485             490             495
Glu Gln Arg Ala Met Leu Ala Gln Gln Tyr Gln Glu Gly Thr Leu Ser
        500             505             510
Val Glu Lys Leu Ala Ala Glu Leu Ser Ala Thr Leu Glu Gln Ala Ala
    515             520             525
Ala Thr Ala Arg Val Thr Phe Ser Gly Leu Arg Asp Thr Val Thr Pro
    530             535             540
Gly Glu Leu Leu Lys Ala Glu Ile Ile Asp Gln Asp Leu Tyr Glu Arg
545             550             555             560
Leu Glu His Gly Gln Ala Thr Ala Lys Asp Val Gly Ser Leu Ala Ser
            565             570             575
Ala Gln Arg Tyr Leu Gln Gly Thr Gly Cys Ile Ala Gly Leu Leu Leu
        580             585             590
Pro Gly Ser Gln Glu Arg Leu Ser Ile Tyr Glu Ala Arg Cys Lys Gly
        595             600             605
Leu Leu Arg Pro Gly Thr Ala Leu Ile Leu Leu Glu Ala Gln Ala Ala
    610             615             620
Thr Gly Phe Ile Ile Asp Pro Lys Ala Asn Lys Gly His Ser Val Glu
625             630             635             640
Glu Ala Leu Arg Ala Ala Val Ile Gly Pro Asp Val Phe Ala Lys Leu
            645             650             655
Leu Ser Ala Glu Arg Ala Val Thr Gly Tyr Thr Asp Pro Tyr Thr Gly
            660             665             670
Gln Gln Ile Ser Leu Phe Gln Ala Met Gln Lys Gly Leu Ile Val Arg
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                 675                      680                      685
Glu His Gly Ile Arg Leu Leu Glu Ala Gln Ile Ala Thr Gly Gly Val
    690                     695                     700
Ile Asp Pro Val His Ser His Arg Val Pro Val Asp Val Ala Tyr Arg
705                     710                     715                 720
Arg Gly Tyr Phe Asp Gln Met Leu Asn Leu Ile Leu Leu Asp Pro Ser
                725                     730                     735
Asp Asp Thr Lys Gly Phe Phe Asp Pro Asn Thr His Glu Asn Leu Thr
            740                     745                     750
Tyr Leu Gln Leu Leu Glu Arg Cys Val Arg Asp Pro Glu Thr Gly Leu
            755                     760                     765
Tyr Leu Leu Pro Leu Ser Ser Thr Gln Ser Pro Leu Val Asp Ser Ala
    770                     775                     780
Thr Gln Gln Ala Phe Gln Asn Leu Leu Leu Ser Val Lys Tyr Gly Arg
785                     790                     795                 800
Phe Gln Gly Gln Arg Val Ser Ala Trp Glu Leu Ile Asn Ser Glu Tyr
                805                     810                     815
Phe Ser Glu Gly Arg Arg Arg Gln Leu Leu Arg Arg Tyr Arg Gln Arg
                820                     825                     830
Glu Val Thr Leu Gly Gln Val Ala Lys Leu Leu Glu Ala Glu Thr Gln
            835                     840                     845
Arg Gln Ala Asp Ile Met Leu Pro Ala Leu Arg Ser Arg Val Thr Val
    850                     855                     860
His Gln Leu Leu Glu Ala Gly Ile Ile Asp Gln Gln Leu Leu Asp Gln
865                     870                     875                 880
Val Leu Ala Gly Thr Ile Ser Pro Glu Ala Leu Leu Leu Met Asp Gly
                885                     890                     895
Val Arg Arg Tyr Leu Cys Gly Leu Gly Ala Val Gly Gly Val Arg Leu
            900                     905                     910
Leu Pro Ser Gly Gln Arg Leu Ser Leu Tyr Gln Ala Met Arg Gln Lys
            915                     920                     925
Leu Leu Gly Pro Arg Val Ala Leu Ala Leu Leu Glu Ala Gln Ala Ala
    930                     935                     940
Thr Gly Thr Ile Met Asp Pro His Ser Pro Glu Ser Leu Ser Val Asp
945                     950                     955                 960
Glu Ala Val Arg Arg Gly Val Val Gly Pro Glu Leu Tyr Gly Arg Leu
                965                     970                     975
Lys Arg Ala Glu Gly Ala Ile Ala Gly Phe Arg Asp Pro Phe Ser Gly
            980                     985                     990
Lys Gln Val Ser Val Phe Gln Ala Met Lys Lys Gly Leu Ile Pro Trp
            995                     1000                    1005
Glu Gln Ala Ala Arg Leu Leu Glu Ala Gln Val Ala Thr Gly Gly Ile
    1010                    1015                    1020
Ile Asp Pro Thr Ser His His His Leu Pro Met Pro Val Ala Ile Gln
1025                    1030                    1035                1040
Arg Gly Tyr Val Asp Gln Glu Met Glu Thr Ala Leu Ser Ser Ser Ser
                1045                    1050                    1055
Glu Thr Phe Pro Thr Pro Asp Gly Gln Gly Arg Thr Ser Tyr Ala Gln
            1060                    1065                    1070
Leu Leu Glu Glu Cys Pro Arg Asp Glu Thr Ser Gly Leu His Leu Leu
            1075                    1080                    1085
Pro Leu Pro Glu Ser Ala Pro Ala Leu Pro Thr Glu Glu Gln Val Gln
    1090                    1095                    1100
Arg Ser Leu Gln Ala Val Pro Gly Ala Lys Asp Gly Thr Ser Leu Trp
1105                    1110                    1115                1120
Asp Leu Leu Ser Ser Cys His Phe Thr Glu Glu Gln Arg Arg Gly Leu
                1125                    1130                    1135
Leu Glu Asp Val Gln Glu Gly Arg Thr Thr Val Pro Gln Leu Leu Ala
            1140                    1145                    1150
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Ser Val Gln Arg Trp Val Gln Glu Thr Lys Leu Leu Ala Gln Ala Arg
        1155   1160   1165
Val Met Val Pro Gly Pro Arg Gly Glu Val Pro Ala Val Trp Leu Leu
        1170   1175   1180
Asp Ala Gly Ile Ile Thr Gln Glu Thr Leu Glu Ala Leu Ala Gln Gly
1185   1190   1195   1200
Thr Gln Ser Pro Ala Gln Val Ala Glu Gln Pro Ala Val Lys Ala Cys
                1205   1210   1215
Leu Trp Gly Thr Gly Cys Val Ala Gly Val Leu Leu Gln Pro Ser Gly
                1220   1225   1230
Ala Lys Ala Ser Ile Ala Gln Ala Val Arg Asp Gly Leu Leu Pro Thr
        1235   1240   1245
Gly Leu Gly Gln Arg Leu Leu Glu Ala Gln Val Ala Ser Gly Phe Leu
        1250   1255   1260
Val Asp Pro Leu Asn Asn Gln Arg Leu Ser Val Glu Asp Ala Val Lys
1265   1270   1275   1280
Val Gly Leu Val Gly Arg Glu Leu Ser Glu Gln Leu Gly Gln Ala Glu
                1285   1290   1295
Arg Ala Ala Ala Gly Tyr Pro Asp Pro Tyr Ser Arg Ala Ser Leu Ser
        1300   1305   1310
Leu Trp Gln Ala Met Glu Lys Gly Leu Val Pro Gln Asn Glu Gly Leu
        1315   1320   1325
Pro Leu Leu Gln Val Gln Leu Ala Thr Gly Gly Val Val Asp Pro Val
        1330   1335   1340
His Gly Val His Leu Pro Gln Ala Ala Ala Cys Arg Leu Gly Leu Leu
1345   1350   1355   1360
Asp Thr Gln Thr Ser Gln Val Leu Thr Ala Val Asp Lys Asp Asn Lys
                1365   1370   1375
Phe Phe Phe Asp Pro Ser Ala Arg Asp Gln Val Thr Tyr Gln Gln Leu
        1380   1385   1390
Arg Glu Arg Cys Val Cys Asp Ser Glu Thr Gly Leu Leu Leu Leu Pro
        1395   1400   1405
Leu Pro Ser Asp Thr Val Leu Glu Val Asp Asp His Thr Ala Val Ala
        1410   1415   1420
Leu Arg Ala Met Lys Val Pro Val Ser Thr Gly Arg Phe Lys Gly Cys
1425   1430   1435   1440
Ser Val Ser Leu Trp Asp Leu Leu Leu Ser Glu Tyr Val Gly Ala Asp
                1445   1450   1455
Lys Arg Arg Glu Leu Val Ala Leu Cys Arg Ser Gly Arg Ala Ala Ala
                1460   1465   1470
Leu Arg Gln Val Val Ser Ala Val Thr Ala Leu Val Glu Ala Ala Glu
        1475   1480   1485
Arg Gln Pro Leu Gln Ala Thr Phe Arg Gly Leu Arg Lys Gln Val Ser
        1490   1495   1500
Ala Arg Asp Leu Phe Arg Ala Gln Leu Ile Ser Arg Lys Thr Leu Asp
1505   1510   1515   1520
Glu Leu Ser Gln Gly Thr Thr Thr Val Lys Glu Val Ala Glu Met Asp
                1525   1530   1535
Ser Val Lys Arg Ser Leu Glu Gly Gly Asn Phe Ile Ala Gly Val Leu
        1540   1545   1550
Ile Gln Gly Thr Gln Glu Arg Met Ser Ile Pro Glu Ala Leu Arg Arg
        1555   1560   1565
His Ile Leu Arg Pro Gly Thr Ala Leu Val Leu Leu Glu Ala Gln Ala
        1570   1575   1580
Ala Thr Gly Phe Ile Ile Asp Pro Ala Glu Asn Arg Lys Leu Thr Val
1585   1590   1595   1600
Glu Glu Ala Phe Lys Ala Gly Met Phe Gly Lys Glu Thr Tyr Val Lys
                1605   1610   1615
Leu Leu Ser Ala Glu Arg Ala Val Thr Gly Tyr Thr Asp Pro Tyr Thr

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              1620                    1625                    1630
Gly Gln Gln Ile Ser Leu Phe Gln Ala Met Gln Lys Asp Leu Ile Val
              1635                    1640                    1645
Arg Glu His Gly Ile Arg Leu Leu Glu Ala Gln Ile Ala Thr Gly Gly
              1650                    1655                    1660
Ile Ile Asp Pro Val His Ser His Arg Val Pro Val Asp Val Ala Tyr
1665                    1670                    1675                    1680
Arg Cys Gly Tyr Phe Asp Glu Glu Met Asn Arg Ile Leu Ala Asp Pro
                   1685                    1690                    1695
Ser Asp Asp Thr Lys Gly Phe Phe Asp Pro Asn Thr His Glu Asn Leu
              1700                    1705                    1710
Thr Tyr Leu Gln Leu Leu Glu Arg Cys Val Glu Asp Pro Glu Thr Gly
              1715                    1720                    1725
Leu Tyr Leu Leu Gln Ile Ile Lys Lys Gly Glu Asn Tyr Val Tyr Ile
              1730                    1735                    1740
Asn Glu Ala Thr Arg His Val Leu Gln Ser Arg Thr Ala Lys Met Arg
1745                    1750                    1755                    1760
Val Gly Arg Phe Ala Asp Gln Val Val Ser Phe Trp Asp Leu Leu Ser
                   1765                    1770                    1775
Ser Pro Tyr Phe Thr Glu Asp Arg Lys Arg Glu Leu Ile Gln Glu Tyr
                   1780                    1785                    1790
Gly Ala Gln Ser Gly Gly Leu Glu Lys Leu Leu Glu Ile Ile Thr Thr
              1795                    1800                    1805
Thr Ile Glu Glu Thr Glu Thr Gln Asn Gln Gly Ile Lys Val Ala Ala
         1810                    1815                    1820
Ile Arg Gly Glu Val Thr Ala Ala Asp Leu Phe Asn Ser Arg Val Ile
1825                    1830                    1835                    1840
Asp Gln Lys Thr Leu His Thr Leu Arg Val Gly Arg Thr Gly Gly Gln
                   1845                    1850                    1855
Ala Leu Ser Thr Leu Glu Cys Val Lys Pro Tyr Leu Glu Gly Ser Asp
              1860                    1865                    1870
Cys Ile Ala Gly Val Thr Val Pro Ser Thr Arg Glu Val Met Ser Leu
         1875                    1880                    1885
His Glu Ala Ser Arg Lys Glu Leu Ile Pro Ala Ala Phe Ala Thr Trp
         1890                    1895                    1900
Leu Leu Glu Ala Gln Ala Ala Thr Gly Phe Leu Leu Asp Pro Cys Thr
1905                    1910                    1915                    1920
Arg Gln Lys Leu Ser Val Asp Glu Ala Val Asp Val Gly Leu Val Asn
                   1925                    1930                    1935
Glu Glu Leu Arg Glu Arg Leu Leu Lys Ala Glu Arg Ala Ala Thr Gly
              1940                    1945                    1950
Tyr Arg Asp Pro Ala Thr Gly Asp Thr Ile Pro Leu Phe Gln Ala Met
         1955                    1960                    1965
Gln Lys Gln Leu Ile Glu Lys Ala Glu Ala Leu Arg Leu Leu Glu Val
         1970                    1975                    1980
Gln Val Ala Thr Gly Gly Val Ile Asp Pro Gln His His His Arg Leu
1985                    1990                    1995                    2000
Pro Leu Glu Thr Ala Tyr Arg Arg Gly Cys Leu His Lys Asp Ile Tyr
                   2005                    2010                    2015
Ala Leu Ile Ser Asp Gln Lys His Met Arg Lys Arg Phe Val Asp Pro
              2020                    2025                    2030
Asn Thr Gln Glu Lys Val Ser Tyr Arg Glu Leu Gln Glu Arg Cys Arg
              2035                    2040                    2045
Pro Gln Glu Asp Thr Gly Trp Val Leu Phe Pro Val Asn Lys Ala Ala
         2050                    2055                    2060
Arg Asp Ser Glu His Ile Asp Asp Glu Thr Arg Arg Ala Leu Glu Ala
2065                    2070                    2075                    2080
Glu Gln Val Glu Ile Thr Val Gly Arg Phe Arg Gly Gln Lys Pro Thr
              2085                    2090                    2095
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102

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Leu Trp Ala Leu Leu Asn Ser Glu Tyr Val Thr Glu Glu Lys Lys Leu
        2100                2105                2110
Gln Leu Val Arg Met Tyr Arg Thr His Thr Arg Arg Ala Leu Gln Thr
        2115                2120                2125
Val Ala Gln Leu Ile Leu Glu Leu Ile Glu Lys Gln Glu Thr Ser Asn
        2130                2135                2140
Lys His Leu Trp Phe Gln Gly Ile Arg Arg Gln Ile Thr Ala Ser Glu
2145                2150                2155                2160
Leu Leu Ser Ser Ala Ile Ile Thr Glu Glu Met Leu Gln Asp Leu Glu
            2165                2170                2175
Thr Gly Arg Ser Thr Thr Gln Glu Leu Met Glu Asp Asp Arg Val Lys
        2180                2185                2190
Arg Tyr Leu Glu Gly Thr Ser Cys Ile Ala Gly Val Leu Val Pro Ala
        2195                2200                2205
Lys Asp Gln Pro Gly Arg Gln Glu Lys Met Ser Ile Tyr Gln Ala Met
        2210                2215                2220
Trp Lys Gly Val Leu Arg Pro Gly Thr Ala Leu Val Leu Leu Glu Ala
2225                2230                2235                2240
Gln Ala Ala Thr Gly Phe Val Ile Asp Pro Val Arg Asn Leu Arg Leu
            2245                2250                2255
Ser Val Glu Glu Pro Val Pro Ala Gly Val Val Gly Ser Glu Ile Gln
        2260                2265                2270
Glu Lys Leu Leu Ser Ala Glu Arg Ala Val Thr Gly Tyr Thr Asp Pro
        2275                2280                2285
Tyr Thr Gly Gln Gln Ile Ser Leu Phe Gln Ala Met Gln Lys Asp Leu
        2290                2295                2300
Ile Val Arg Glu His Gly Ile Arg Leu Leu Glu Ala Gln Ile Ala Thr
2305                2310                2315                2320
Gly Gly Val Ile Asp Pro Val His Ser His Arg Val Pro Val Asp Val
            2325                2330                2335
Ala Tyr Arg Arg Gly Tyr Phe Asp Glu Glu Met Asn Arg Val Leu Ala
        2340                2345                2350
Asp Pro Ser Asp Asp Thr Lys Gly Phe Phe Asp Pro Asn Thr His Glu
        2355                2360                2365
Asn Leu Thr Tyr Val Gln Leu Leu Arg Arg Cys Val Pro Asp Pro Asp
        2370                2375                2380
Thr Gly Leu Tyr Met Leu Gln Leu Ala Gly Arg Gly Ser Ala Val His
2385                2390                2395                2400
Gln Leu Ser Glu Glu Leu Arg Cys Ala Leu Arg Asp Ala Arg Val Thr
            2405                2410                2415
Pro Gly Ser Gly Ala Leu Gln Gly Gln Ser Val Ser Val Trp Glu Leu
        2420                2425                2430
Leu Phe Tyr Arg Glu Val Ser Glu Asp Arg Arg Gln Asp Leu Leu Ser
        2435                2440                2445
Arg Tyr Arg Ala Gly Thr Leu Thr Val Glu Glu Leu Gly Ala Thr Leu
        2450                2455                2460
Thr Ser Leu Leu Ala Gln Ala Gln Ala Gln Ala Arg Ala Glu Ala Glu
2465                2470                2475                2480
Ala Gly Ser Pro Arg Pro Asp Pro Arg Glu Ala Leu Arg Ala Ala Thr
            2485                2490                2495
Met Glu Val Lys Val Gly Arg Leu Arg Gly Arg Ala Val Pro Val Trp
        2500                2505                2510
Asp Val Leu Ala Ser Gly Tyr Val Ser Arg Ala Ala Arg Glu Glu Leu
        2515                2520                2525
Leu Ala Glu Phe Gly Ser Gly Thr Leu Asp Leu Pro Ala Leu Thr Arg
        2530                2535                2540
Arg Leu Thr Ala Ile Ile Glu Glu Ala Glu Glu Ala Pro Gly Ala Arg
2545                2550                2555                2560
Pro Gln Leu Gln Asp Ala Arg Arg Gly Pro Arg Glu Pro Gly Pro Ala
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                  2565                2570                2575
Gly Arg Gly Asp Gly Asp Ser Gly Arg Ser Gln Arg Glu Gly Gln Gly
              2580                2585                2590
Glu Gly Glu Thr Gln Glu Ala Ala Ala Ala Ala Ala Ala Ala Arg Arg
              2595                2600                2605
Gln Glu Gln Thr Leu Arg Asp Ala Thr Met Glu Val Gln Arg Gly Gln
              2610                2615                2620
Phe Gln Gly Arg Pro Val Ser Val Trp Asp Val Leu Phe Ser Ser Tyr
2625                2630                2635                2640
Leu Ser Glu Ala Arg Arg Asp Glu Leu Leu Ala Gln His Ala Ala Gly
              2645                2650                2655
Ala Leu Gly Leu Pro Asp Leu Val Ala Val Leu Thr Arg Val Ile Glu
              2660                2665                2670
Glu Thr Glu Glu Arg Leu Ser Lys Val Ser Phe Arg Gly Leu Arg Arg
              2675                2680                2685
Gln Val Ser Ala Ser Glu Leu His Thr Ser Gly Ile Leu Gly Pro Glu
              2690                2695                2700
Thr Leu Arg Asp Leu Ala Gln Gly Thr Lys Thr Leu Gln Glu Val Thr
2705                2710                2715                2720
Glu Met Asp Ser Val Lys Arg Tyr Leu Glu Gly Thr Ser Cys Ile Ala
              2725                2730                2735
Gly Val Leu Val Pro Ala Lys Asp Gln Pro Gly Arg Gln Glu Lys Met
              2740                2745                2750
Ser Ile Tyr Gln Ala Met Trp Lys Gly Val Leu Arg Pro Gly Thr Ala
              2755                2760                2765
Leu Val Leu Leu Glu Ala Gln Ala Ala Thr Gly Phe Val Ile Asp Pro
              2770                2775                2780
Val Arg Asn Leu Arg Leu Ser Val Glu Glu Ala Val Ala Ala Gly Val
2785                2790                2795                2800
Val Gly Gly Glu Ile Gln Glu Lys Leu Leu Ser Ala Glu Arg Ala Val
              2805                2810                2815
Thr Gly Tyr Thr Asp Pro Tyr Thr Gly Gln Gln Ile Ser Leu Phe Gln
              2820                2825                2830
Ala Met Gln Lys Asp Leu Ile Val Arg Glu His Gly Ile Arg Leu Leu
              2835                2840                2845
Glu Ala Gln Ile Ala Thr Gly Gly Val Ile Asp Pro Val His Ser His
              2850                2855                2860
Arg Val Pro Val Asp Val Ala Tyr Arg Arg Gly Tyr Phe Asp Glu Glu
2865                2870                2875                2880
Met Asn Arg Val Leu Ala Asp Pro Ser Asp Asp Thr Lys Gly Phe Phe
              2885                2890                2895
Asp Pro Asn Thr His Glu Asn Leu Thr Tyr Val Gln Leu Leu Arg Arg
              2900                2905                2910
Cys Val Pro Asp Pro Asp Thr Gly Leu Tyr Met Leu Gln Leu Ala Gly
              2915                2920                2925
Arg Gly Ser Ala Val His Gln Leu Ser Glu Glu Leu Arg Cys Ala Leu
              2930                2935                2940
Arg Asp Ala Arg Val Thr Pro Gly Ser Gly Ala Leu Gln Gly Gln Ser
2945                2950                2955                2960
Val Ser Val Trp Glu Leu Leu Phe Tyr Arg Glu Val Ser Glu Asp Arg
              2965                2970                2975
Arg Gln Asp Leu Leu Ser Arg Tyr Arg Ala Gly Thr Leu Thr Val Glu
              2980                2985                2990
Glu Leu Gly Ala Thr Leu Thr Ser Leu Leu Ala Gln Ala Gln Ala Gln
              2995                3000                3005
Ala Arg Ala Glu Ala Glu Ala Gly Ser Pro Arg Pro Asp Pro Arg Glu
              3010                3015                3020
Ala Leu Arg Ala Ala Thr Met Glu Val Lys Val Gly Arg Leu Arg Gly
3025                3030                3035                3040
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Arg Ala Val Pro Val Trp Asp Val Leu Ala Ser Gly Tyr Val Ser Gly
              3045                3050                3055
Ala Ala Arg Glu Glu Leu Leu Ala Glu Phe Gly Ser Gly Thr Leu Asp
              3060                3065                3070
Leu Pro Ala Leu Thr Arg Arg Leu Thr Ala Ile Ile Glu Glu Ala Glu
              3075                3080                3085
Glu Ala Pro Gly Ala Arg Pro Gln Leu Gln Asp Ala Trp Arg Gly Pro
              3090                3095                3100
Arg Glu Pro Gly Pro Ala Gly Arg Gly Asp Gly Asp Ser Gly Arg Ser
3105                3110                3115                3120
Gln Arg Glu Gly Gln Gly Glu Gly Glu Thr Gln Glu Ala Ala Ala Ala
              3125                3130                3135
Ala Ala Ala Ala Arg Arg Gln Glu Gln Thr Leu Arg Asp Ala Thr Met
              3140                3145                3150
Glu Val Gln Arg Gly Gln Phe Gln Gly Arg Pro Val Ser Val Trp Asp
              3155                3160                3165
Val Leu Phe Ser Ser Tyr Leu Ser Glu Ala Arg Arg Asp Glu Leu Leu
              3170                3175                3180
Ala Gln His Ala Ala Gly Ala Leu Gly Leu Pro Asp Leu Val Ala Val
3185                3190                3195                3200
Leu Thr Arg Val Ile Glu Glu Thr Glu Glu Arg Leu Ser Lys Val Ser
              3205                3210                3215
Phe Arg Gly Leu Arg Arg Gln Val Ser Ala Ser Glu Leu His Thr Ser
              3220                3225                3230
Gly Ile Leu Gly Pro Glu Thr Leu Arg Asp Leu Ala Gln Gly Thr Lys
              3235                3240                3245
Thr Leu Gln Glu Val Thr Glu Met Asp Ser Val Lys Arg Tyr Leu Glu
              3250                3255                3260
Gly Thr Ser Cys Ile Ala Gly Val Leu Val Pro Ala Lys Asp Gln Pro
3265                3270                3275                3280
Gly Arg Gln Glu Lys Met Ser Ile Tyr Gln Ala Met Trp Lys Gly Val
              3285                3290                3295
Leu Arg Pro Gly Thr Ala Leu Val Leu Leu Glu Ala Gln Ala Ala Thr
              3300                3305                3310
Gly Phe Val Ile Asp Pro Val Arg Asn Leu Arg Leu Ser Val Glu Glu
              3315                3320                3325
Ala Val Ala Ala Gly Val Val Gly Gly Glu Ile Gln Glu Lys Leu Leu
3330                3335                3340
Ser Ala Glu Arg Ala Val Thr Gly Tyr Thr Asp Pro Tyr Thr Gly Gln
3345                3350                3355                3360
Gln Ile Ser Leu Phe Gln Ala Met Gln Lys Asp Leu Ile Val Arg Glu
              3365                3370                3375
His Gly Ile Arg Leu Leu Glu Ala Gln Ile Ala Thr Gly Gly Val Ile
              3380                3385                3390
Asp Pro Val His Ser His Arg Val Pro Val Asp Val Ala Tyr Arg Arg
              3395                3400                3405
Gly Tyr Phe Asp Glu Glu Met Asn Arg Val Leu Ala Asp Pro Ser Asp
              3410                3415                3420
Asp Thr Lys Gly Phe Phe Asp Pro Asn Thr His Glu Asn Leu Thr Tyr
3425                3430                3435                3440
Val Gln Leu Leu Arg Arg Cys Val Pro Asp Pro Asp Thr Gly Leu Tyr
              3445                3450                3455
Met Leu Gln Leu Ala Gly Arg Gly Ser Ala Val His Gln Leu Ser Glu
              3460                3465                3470
Glu Leu Arg Cys Ala Leu Arg Asp Ala Arg Val Thr Pro Gly Ser Gly
3475                3480                3485
Ala Leu Gln Gly Gln Ser Val Ser Val Trp Glu Leu Leu Phe Tyr Arg
              3490                3495                3500
Glu Val Ser Glu Asp Arg Arg Gln Asp Leu Leu Ser Arg Tyr Arg Ala
```

```
3505                 3510                 3515                 3520
Gly Thr Leu Thr Val Glu Glu Leu Gly Ala Thr Leu Thr Ser Leu Leu
            3525                 3530                 3535
Ala Gln Ala Gln Ala Gln Ala Arg Ala Glu Ala Glu Ala Gly Ser Pro
            3540                 3545                 3550
Arg Pro Asp Pro Arg Glu Ala Leu Arg Ala Ala Thr Met Glu Val Lys
            3555                 3560                 3565
Val Gly Arg Leu Arg Gly Arg Ala Val Pro Val Trp Asp Val Leu Ala
            3570                 3575                 3580
Ser Gly Tyr Val Ser Gly Ala Ala Arg Glu Glu Leu Leu Ala Glu Phe
3585                 3590                 3595                 3600
Gly Ser Gly Thr Leu Asp Leu Pro Ala Leu Thr Arg Arg Leu Thr Ala
            3605                 3610                 3615
Ile Ile Glu Glu Ala Glu Glu Ala Pro Gly Ala Arg Pro Gln Leu Gln
            3620                 3625                 3630
Asp Ala Trp Arg Gly Pro Arg Glu Pro Gly Pro Ala Gly Arg Gly Asp
            3635                 3640                 3645
Gly Asp Ser Gly Arg Ser Gln Arg Glu Gly Gln Gly Glu Gly Glu Thr
            3650                 3655                 3660
Gln Glu Ala Ala Ala Ala Ala Ala Ala Arg Arg Gln Glu Gln Thr
3665                 3670                 3675                 3680
Leu Arg Asp Ala Thr Met Glu Val Gln Arg Gly Gln Phe Gln Gly Arg
            3685                 3690                 3695
Pro Val Ser Val Trp Asp Val Leu Phe Ser Ser Tyr Leu Ser Glu Ala
            3700                 3705                 3710
Arg Arg Asp Glu Leu Leu Ala Gln His Ala Ala Gly Ala Leu Gly Leu
            3715                 3720                 3725
Pro Asp Leu Val Ala Val Leu Thr Arg Val Ile Glu Glu Thr Glu Glu
            3730                 3735                 3740
Arg Leu Ser Lys Val Ser Phe Arg Gly Leu Arg Arg Gln Val Ser Ala
3745                 3750                 3755                 3760
Ser Glu Leu His Thr Ser Gly Ile Leu Gly Pro Glu Thr Leu Arg Asp
            3765                 3770                 3775
Leu Ala Gln Gly Thr Lys Thr Leu Gln Glu Val Thr Glu Met Asp Ser
            3780                 3785                 3790
Val Lys Arg Tyr Leu Glu Gly Thr Ser Cys Ile Ala Gly Val Leu Val
            3795                 3800                 3805
Pro Ala Lys Asp Gln Pro Gly Arg Gln Glu Lys Met Ser Ile Tyr Gln
            3810                 3815                 3820
Ala Met Trp Lys Gly Val Leu Arg Pro Gly Thr Ala Leu Val Leu Leu
3825                 3830                 3835                 3840
Glu Ala Gln Ala Ala Thr Gly Phe Val Ile Asp Pro Val Arg Asn Leu
            3845                 3850                 3855
Arg Leu Ser Val Glu Glu Ala Val Ala Ala Gly Val Val Gly Gly Glu
            3860                 3865                 3870
Ile Gln Glu Lys Leu Leu Ser Ala Glu Arg Ala Val Thr Gly Tyr Thr
            3875                 3880                 3885
Asp Pro Tyr Thr Gly Gln Gln Ile Ser Leu Phe Gln Ala Met Gln Lys
            3890                 3895                 -   3900
Asp Leu Ile Val Arg Glu His Gly Ile Arg Leu Leu Glu Ala Gln Ile
3905                 3910                 3915                 3920
Ala Thr Gly Gly Val Ile Asp Pro Val His Ser His Arg Val Pro Val
            3925                 3930                 3935
Asp Val Ala Tyr Arg Arg Gly Tyr Phe Asp Glu Glu Met Asn Arg Val
            3940                 3945                 3950
Leu Ala Asp Pro Ser Asp Asp Thr Lys Gly Phe Phe Asp Pro Asn Thr
            3955                 3960                 3965
His Glu Asn Leu Thr Tyr Val Gln Leu Leu Arg Arg Cys Val Pro Asp
            3970                 3975                 3980
```

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Pro Asp Thr Gly Leu Tyr Met Leu Gln Leu Ala Gly Arg Gly Ser Ala
3985                3990                3995                4000
Val His Gln Leu Ser Glu Glu Leu Arg Cys Ala Leu Arg Asp Ala Arg
                4005                4010                4015
Val Thr Pro Gly Ser Gly Ala Leu Gln Gly Gln Ser Val Ser Val Trp
            4020                4025                4030
Glu Leu Leu Phe Tyr Arg Glu Val Ser Glu Asp Arg Arg Gln Asp Leu
            4035                4040                4045
Leu Ser Arg Tyr Arg Ala Ser Thr Leu Thr Val Glu Glu Leu Gly Ala
        4050                4055                4060
Thr Leu Thr Ser Leu Leu Ala Gln Ala Gln Ala Gln Ala Arg Ala Glu
4065                4070                4075                4080
Ala Glu Ala Gly Ser Pro Arg Pro Asp Pro Arg Glu Ala Leu Arg Ala
                4085                4090                4095
Ala Thr Met Glu Val Lys Val Gly Arg Leu Arg Gly Arg Ala Val Pro
            4100                4105                4110
Val Trp Asp Val Leu Ala Ser Gly Tyr Val Ser Arg Ala Ala Arg Glu
            4115                4120                4125
Glu Leu Leu Ala Glu Phe Gly Ser Gly Thr Leu Asp Leu Pro Ala Leu
        4130                4135                4140
Thr Arg Arg Leu Thr Ala Ile Ile Glu Glu Ala Glu Glu Ala Pro Gly
4145                4150                4155                4160
Ala Arg Pro Gln Leu Gln Asp Ala Trp Arg Gly Pro Arg Glu Pro Gly
            4165                4170                4175
Pro Ala Gly Arg Gly Asp Gly Asp Ser Gly Arg Ser Gln Arg Glu Gly
            4180                4185                4190
Gln Gly Glu Gly Glu Thr Gln Glu Ala Ala Ala Ala Thr Ala Ala Ala
        4195                4200                4205
Arg Arg Gln Glu Gln Thr Leu Arg Asp Ala Thr Met Glu Val Gln Arg
        4210                4215                4220
Gly Gln Phe Gln Gly Arg Pro Val Ser Val Trp Asp Val Leu Phe Ser
4225                4230                4235                4240
Ser Tyr Leu Ser Glu Ala Arg Arg Asp Gly Leu Leu Ala Gln His Ala
            4245                4250                4255
Ala Gly Ala Leu Gly Leu Pro Asp Leu Val Ala Val Leu Thr Arg Val
        4260                4265                4270
Ile Glu Glu Thr Glu Glu Arg Leu Ser Lys Val Ser Phe Arg Gly Leu
    4275                4280                4285
Arg Arg Gln Val Ser Ala Ser Glu Leu His Thr Ser Gly Ile Leu Gly
    4290                4295                4300
Pro Glu Thr Leu Arg Asp Leu Ala Gln Gly Thr Lys Thr Leu Gln Glu
4305                4310                4315                4320
Val Thr Glu Met Asp Ser Val Lys Arg Tyr Leu Glu Gly Thr Ser Cys
            4325                4330                4335
Ile Ala Gly Val Leu Val Pro Ala Lys Asp Gln Pro Gly Arg Gln Glu
            4340                4345                4350
Lys Met Ser Ile Tyr Gln Ala Met Trp Lys Gly Val Leu Arg Pro Gly
            4355                4360                4365
Thr Ala Leu Val Leu Leu Glu Ala Gln Ala Ala Thr Gly Phe Val Ile
    4370                4375                4380
Asp Pro Val Arg Asn Leu Arg Leu Ser Val Glu Glu Ala Val Ala Ala
4385                4390                4395                4400
Gly Val Val Gly Gly Glu Ile Gln Glu Lys Leu Leu Ser Ala Glu Arg
            4405                4410                4415
Ala Val Thr Gly Tyr Thr Asp Pro Tyr Thr Gly Gln Gln Ile Ser Leu
            4420                4425                4430
Phe Gln Ala Met Gln Lys Asp Leu Ile Val Arg Glu His Gly Ile Arg
        4435                4440                4445
Leu Leu Glu Ala Gln Ile Ala Thr Gly Gly Val Ile Asp Pro Val His
```

```
        4450              4455              4460
Ser His Arg Val Pro Val Asp Val Ala Tyr Arg Arg Gly Tyr Phe Asp
4465              4470              4475              4480
Glu Glu Met Asn Arg Val Leu Ala Asp Pro Ser Asp Asp Thr Lys Gly
                  4485              4490              4495
Phe Phe Asp Pro Asn Thr His Glu Asn Leu Thr Tyr Val Gln Leu Leu
              4500              4505              4510
Arg Arg Cys Val Pro Asp Pro Asp Thr Gly Leu Tyr Met Leu Gln Leu
              4515              4520     ·        4525
Ala Gly Arg Gly Ser Ala Val His Gln Leu Ser Glu Glu Leu Arg Cys
          4530              4535.             4540
Ala Leu Arg Asp Ala Arg Val Thr Pro Gly Ser Gly Ala Leu Gln Gly
4545              4550     ·        4555              4560
Gln Ser Val Ser Val Trp Glu Leu Leu Phe Tyr Arg Glu Val Ser Glu
                  4565              4570              4575
Asp Arg Arg Gln Asp Leu Leu Ser Arg Tyr Arg Ala Gly Thr Leu Thr
                  4580              4585 ·            4590
Val Glu Glu Leu Gly Ala Thr Leu Thr Ser Leu Leu Ala Gln Ala Gln
                  4595              4600              4605
Ala Gln Ala Arg Ala Glu Ala Glu Ala Gly Ser Pro Arg Pro Asp Pro
          4610              4615              4620
Arg Glu Ala Leu Arg Ala Ala Thr Met Glu Val Lys Val Gly Arg Leu
4625              4630              4635              4640
Arg Gly Arg Ala Val Pro Val Trp Asp Val Leu Ala Ser Gly Tyr Val
                  4645              4650              4655
Ser Gly Ala Ala Arg Glu Glu Leu Leu Ala Glu Phe Gly Ser Gly Thr
                  4660              4665              4670
Leu Asp Leu Pro Ala Leu Thr Arg Arg Leu Thr Ala Ile Ile Glu Glu
                  4675              4680              4685
Ala Glu Glu Ala Pro Gly Ala Arg Pro Gln Leu Gln Asp Ala Trp Arg
          .4690              4695              4700
Gly Pro Arg Glu Pro Gly Pro Ala Gly Arg Gly Asp Gly Asp Ser Gly
4705              4710              4715              4720
Arg Ser Gln Arg Glu Gly Gln Gly Glu Gly Glu Thr Gln Glu Ala Ala
                  4725              4730              4735
Ala Ala Ala Ala Ala Ala Arg Arg Gln Glu Gln Thr Leu Arg Asp Ala
          4740              4745              4750
Thr Met Glu Val Gln Arg Gly Gln Phe Gln Gly Arg Pro Val Ser Val
          4755              4760              4765
Trp Asp Val Leu Phe Ser Ser Tyr Leu Ser Glu Ala Arg Arg Asp Glu
          4770              4775              4780
Leu Leu Ala Gln His Ala Ala Gly Ala Leu Gly Leu Pro Asp Leu Val
4785              4790              4795              4800
Ala Val Leu Thr Arg Val Ile Glu Glu Thr Glu Glu Arg Leu Ser Lys
          4805              4810              4815
Val Ser Phe Arg Gly Leu Arg Arg Gln Val Ser Ala Ser Glu Leu His
          4820              4825              4830
Thr Ser Gly Ile Leu Gly Pro Glu Thr Leu Arg Asp Leu Ala Gln Gly
          4835              4840              4845
Thr Lys Thr Leu Gln Glu Val Thr Glu Met Asp Ser Val Lys Arg Tyr
          4850              4855              4860
Leu Glu Gly Thr Ser Cys Ile Ala Gly Val Leu Val Pro Ala Lys Asp
4865              4870              4875              4880
Gln Pro Gly Arg Gln Glu Lys Met Ser Ile Tyr Gln Ala Met Trp Lys
                  4885              4890              4895
Gly Val Leu Arg Pro Gly Thr Ala Leu Val Leu Leu Glu Ala Gln Ala
                  4900              4905              4910
Ala Thr Gly Phe Val Ile Asp Pro Val Arg Asn Leu Arg Leu Ser Val
          4915              4920              4925
```

```
Glu Glu Ala Val Ala Ala Gly Val Val Gly Gly Glu Ile Gln Glu Lys
    4930             4935             4940
Leu Leu Ser Ala Glu Arg Ala Val Thr Gly Tyr Thr Asp Pro Tyr Thr
4945             4950             4955             4960
Gly Gln Gln Ile Ser Leu Phe Gln Ala Met Gln Lys Asp Leu Ile Val
            4965             4970             4975
Arg Glu His Gly Ile Arg Leu Leu Glu Ala Gln Ile Ala Thr Gly Gly
            4980             4985             4990
Val Ile Asp Pro Val His Ser His Arg Val Pro Val Asp Val Ala Tyr
            4995         5000             5005
Arg Arg Gly Tyr Phe Asp Glu Glu Met Asn Arg Val Leu Ala Asp Pro
    5010             5015             5020
Ser Asp Asp Thr Lys Gly Phe Phe Asp Pro Asn Thr His Glu Asn Leu
5025             5030             5035             5040
Thr Tyr Leu Gln Leu Leu Gln Arg Ala Thr Leu Asp Pro Glu Thr Gly
            5045             5050             5055
Leu Leu Phe Leu Ser Leu Ser Leu Gln
            5060             5065
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```
<210> 53
<211> 1664
<212> DNA
<213> Homo sapiens

<400> 53
tcatggccgg ctcctaccct gaaggtgcac ctgcaatcct cgccgataag aggcagcagt 60
tcggaagccg gttcctgagc gatccggcgc gggtcttcca ccacaatgcc tgttgattat 120
gagatcaatg cccacaaata ctggaatgac ttctacaaaa tccacgaaaa tgggttttc 180
aaggatagac attggctttt taccgaattc cctgagctgg cacctagcca aatcaaaat 240
catttgaagg attggttctt ggagaacaag agtgaagtat gtgaatgtag aaacaatgag 300
gatggacctg gtttaataat ggaagaacag cacaagtgtt cttcgaagag ccttgaacat 360
aaaacacaga cacctcctgt ggaggagaat gtaactcaga aaattagtga cctggaaatt 420
tgtgctgatg agtttcctgg atcctcagcc acctaccgaa tactggaggt tggctgtggt 480
gtgggaaaca cagtctttcc aatttacaa acgaacaatg acccaggact ctttgtttat 540
tgctgtgatt tttcttccac agctatagaa ctggtccaga caaattcaga atatgatcct 600
tctcggtgtt ttgcctttgt tcacgacctg tgtgatgaag agaagagtta cccagtgccc 660
aagggcagtc ttgatattat cattctcata tttgttcttt cagcaattgt tccagacaag 720
atgcagaagg ctatcaacag gctgagcagg cttctgaaac ctggggggat ggtacttctg 780
cgagattacg gccgctatga catggctcag cttcggttta aaaaaggtca gtgtctatct 840
ggaaatttct atgtgagagg tgatggaacc agagtttact tcttcacaca agaggaactg 900
gacacgcttt tcaccactgc tggactggaa aaagttcaga acctggtgga ccgccgactg 960
caggtgaacc gagggaagca actgacaatg taccgggttt ggattcagtg caaatactgc 1020
aagccccttc tgtccagcac cagctaagag gcacctgctg ccaacacgat gcaagcccgt 1080
tgtgtttccg agcttttttt aaaaaaaaat ttgtagcacc gggcatggtg catgcctgta 1140
atcccagcca ctcaggaggc tgaggcaggg aggatccatt gagcccagga gtccagcctg 1200
ggcaaaatag cgagagaccc tgaatctgaa agtaatgata aaataaaaag aatataaatg 1260
aggtctcgtt gatgctggac aattcaagaa ttcagacttg aaccttaaac ctaggaaaag 1320
ttactttgta tcaggattct aacaattatg cttcatattt gtgaagtcct ttaaaacata 1380
attttctcaa gttctttctt tgagacctca atctgtctta gcattttgta actaataact 1440
gaaattttat tcaaaggaat tgtaaacctt aaaccaccaa tttatttcca tgtgaaaaag 1500
tgttatatat gacaagtgtt ttttgattgt aattgcgtta aatcttttga gagtgtaaat 1560
gccgggctag gcaattgcag ttaatacata caggggttag tgaagggctt attaagttgt 1620
aggggaagca agctgggaag aatcagatca gatattttcc tgac                 1664

<210> 54
<211> 313
<212> PRT
<213> Homo sapiens
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<400> 54
Met Pro Val Asp Tyr Glu Ile Asn Ala His Lys Tyr Trp Asn Asp Phe
 1               5                  10              15
Tyr Lys Ile His Glu Asn Gly Phe Phe Lys Asp Arg His Trp Leu Phe
             20              25              30
Thr Glu Phe Pro Glu Leu Ala Pro Ser Gln Asn Gln Asn His Leu Lys
         35              40              45
Asp Trp Phe Leu Glu Asn Lys Ser Glu Val Cys Glu Cys Arg Asn Asn
     50              55              60
Glu Asp Gly Pro Gly Leu Ile Met Glu Glu Gln His Lys Cys Ser Ser
65              70              75              80
Lys Ser Leu Glu His Lys Thr Gln Thr Pro Pro Val Glu Glu Asn Val
             85              90              95
Thr Gln Lys Ile Ser Asp Leu Glu Ile Cys Ala Asp Glu Phe Pro Gly
         100             105             110
Ser Ser Ala Thr Tyr Arg Ile Leu Glu Val Gly Cys Gly Val Gly Asn
         115             120             125
Thr Val Phe Pro Ile Leu Gln Thr Asn Asn Asp Pro Gly Leu Phe Val
     130             135             140
Tyr Cys Cys Asp Phe Ser Ser Thr Ala Ile Glu Leu Val Gln Thr Asn
145             150             155             160
Ser Glu Tyr Asp Pro Ser Arg Cys Phe Ala Phe Val His Asp Leu Cys
             165             170             175
Asp Glu Glu Lys Ser Tyr Pro Val Pro Lys Gly Ser Leu Asp Ile Ile
             180             185             190
Ile Leu Ile Phe Val Leu Ser Ala Ile Val Pro Asp Lys Met Gln Lys
         195             200             205
Ala Ile Asn Arg Leu Ser Arg Leu Leu Lys Pro Gly Gly Met Val Leu
     210             215             220
Leu Arg Asp Tyr Gly Arg Tyr Asp Met Ala Gln Leu Arg Phe Lys Lys
.225             230             235             240
Gly Gln Cys Leu Ser Gly Asn Phe Tyr Val Arg Gly Asp Gly Thr Arg
             245             250             255
Val Tyr Phe Phe Thr Gln Glu Glu Leu Asp Thr Leu Phe Thr Thr Ala
         260             265             270
Gly Leu Glu Lys Val Gln Asn Leu Val Asp Arg Arg Leu Gln Val Asn
     275             280             285
Arg Gly Lys Gln Leu Thr Met Tyr Arg Val Trp Ile Gln Cys Lys Tyr
     290             295             300
Cys Lys Pro Leu Leu Ser Ser Thr Ser
305             310
```

```
<210> 55
<211> 3334
<212> DNA
<213> Homo sapiens
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<400> 55
gaaaaggaaa tcgcagctgt gatttctcct gaactggagc atctagataa aacccttccc 60
accatgaata atctcatcag ccaagataag cgtatcagct ctaaccctgt ggccaaaata 120
atatatggtg acccagtgac cttcctgccc cacctgcccc ggaaaagtgt ggtccattgc 180
tctaagattt ggagctgcag gaaaagaatt acagttgagt acctccagca cattgtggaa 240
cagaaaaatg gcaaagaaag agtgcccatc ctctggcatt tcctgcagaa ggaagcagag 300
ctgaggctgg taaagttcct gcctgagatt ttggccttgc aaagggatct agtgaagcag 360
ttccagaacg ttcagcaagt tgaatacagc tccatcagag gcttcctcag caagcacagc 420
tcagatgggt tgaggcagct gcttcacaac aggatcacag tctttctgtc cacatggaac 480
aaactgagga gatcgcttga gacgaacggt gagatcaacc tacccaaaga ctactgcagc 540
```

```
actgacttgg atctggacac tgagtttgag atcctcttgc cacgccgacg gggcctgggc 600
ctctgtgcta ccgctctcgt cagctacttg attcgcctac acaatgaaat tgtctacgcc 660
gtggaaaaac tctccaagga aaacaacagc tattccgtgg atgccgccga ggtcactgaa 720
ctgcatgtca tcagttatga agtggagcgg gacctgactc cactgattct ctccaactgc 780
cagtaccagg tggaggaggg cagagagacc gtgcaggagt tcgatctgga gaagattcag 840
cggcagatcg tcagccgctt cctccagggc aagccccggc tgagcctcaa gggaataccc 900
actctggtgt acagacacga ctggaactat gaacatctct ttatggacat caagaacaaa 960
atggcacagg actccctccc cagctcggtc attagtgcca tcagtggaca gctgcagtcc 1020
tacagcgatg cctgtgaagt gctgtctgtc gtagaagtca ctctgggggtt tctgagcaca 1080
gctggtgggg atccaaacat gcagctgaat gtgtatactc aagacatcct gcaaatgggt 1140
gatcagacga ttcacgtgtt aaaggcctta aacagatgcc agttaaaaca caccattgcc 1200
ctctggcagt tcctgtctgc tcataagtct gaacagctgc tgcggctgca caaagagcca 1260
tttgggggaaa tcagttcaag gtacaaagcg gatctgagcc cggaaaatgc taagctcctc 1320
agcacattcc taaatcagac tggcctagac gccttcctgc tagagctgca cgaaatgata 1380
atcttgaaac taaagaaccc ccaaacccaa accgaggagc gcttccgccc tcagtggagc 1440
ctgagagaca ctctcgtaag ttacatgcaa actaaagaaa gtgaaattct tcctgaaatg 1500
gcatctcagt tcccagaaga gatactgctc gccagctgtg tctcagtgtg gaaaacagct 1560
gctgtgctga aatggaatcg agaaatgaga tagaattatt tcctcagcta tctttggatg 1620
actttggaga gaagactcct ctctcctcgt ctgcggcgtg gacttgatca tggactggtg 1680
cctttgcatt cagaaggaga gctgtcagcg tagcaccgaa ttcaagacca aggcgtgcta 1740
cctgagctga cagctttttg aaagccgagc tgtttctgaa ccatgtacat acatgttctg 1800
aaactttctc atcattttat gagtactgtt cattgagaga tgacaatgaa gattagatga 1860
aattggaaat aaaccaacat tgtttacatt ccaggagact tgtagctcag ccacacacgc 1920
agtaatgacc tgtgcccgtt cgcctctggc actgcccacc cctctttttt tttttcttct 1980
aattctgtac tcacaaaaga gaatctcatt ttcttctttc ttccattccc ttaaattctg 2040
agtactgtac atatatttct gggttcccac gatgatgtga aaaactacca gactgttttt 2100
tgtcttctca caaagacaag aaaaatcagg gcattttgtg agtgccttaa gatcaaacta 2160
acaagatctg accctctccc ctcacagtga gccactgccc cacttcagag ggtaagagcc 2220
aaaagcctca ttgtgaaagg cactggactt ggaccaggga caccatcagg gccttggttt 2280
tctcacgcat aaaatggaga gtggattaat cgccaaagat tcttctgatc tgacattttg 2340
aaattgtgag agaaactaga tgactgtaaa cttggtcaca ggcctggttc tggcagttct 2400
ttgcggactt ttttctagca ttatgccaaa taaacatgca gtctcagtgt gctctcgcat 2460
gtatgaatat ctagtccttt ctgtggttct cagccaagac ataaaaacta ggactcagag 2520
cacatacaaa accagttatg tttcggaaag agggaaaaga gtccccgagc ccggatcttg 2580
tgctgctttt ctcactacg tgttgccttt tttctttaca aaatctgctt tgatacttag 2640
gacctctctg gactaatttc tcttcctaga cagctcagca cagctcattga tatgttagag 2700
gcagtatcct taatattcat tctaaatgag ttaacgactt aacttgaaat tgggcctaag 2760
gagtgagaac tacaaaaata caaaatgctt gtccaggact cagccatgca caccttgagc 2820
agcgccggca ggaggcacgg aaggaactgt gctccgttct cctcactgtc atggtgccac 2880
cagtgtctga tgaagggcag agtgacccag actgcaggca gtaactgact tcacacagtc 2940
cctggcattt agtcatctgt gattgtttta tcactctgga ctgtgcagag ccacctgcca 3000
ccgagatctg cattccgact gcctatgaac gggtgtgggg gccggggggct ggcttgctga 3060
agtcttcaac ttgcactcgg agctcctttg atacctcaga gctggctgtc aggtggcagc 3120
tcacacccag actcactggc cacacctcag cagggggggga gtcgagtgtc agtctctttc 3180
tgtgaaggct ttttttttcc tttggcctgg gaattttttcc cattttttatg aaggggtttt 3240
aaattgtttc attttgtgtg ctgtgcttca aagccttaac tgtcaaatct tgcattatct 3300
tgtttgtaca gaaatatact ggcctagcag aggc 3334
```

<210> 56
<211> 509
<212> PRT
<213> Homo sapiens

<400> 56

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Met Asn Asn Leu Ile Ser Gln Asp Lys Arg Ile Ser Ser Asn Pro Val
1               5                   10                  15
Ala Lys Ile Ile Tyr Gly Asp Pro Val Thr Phe Leu Pro His Leu Pro
            20                  25                  30
Arg Lys Ser Val Val His Cys Ser Lys Ile Trp Ser Cys Arg Lys Arg
```

```
            35                    40                    45
Ile Thr Val Glu Tyr Leu Gln His Ile Val Glu Gln Lys Asn Gly Lys
    50                  55                  60
Glu Arg Val Pro Ile Leu Trp His Phe Leu Gln Lys Glu Ala Glu Leu
65                  70                  75                  80
Arg Leu Val Lys Phe Leu Pro Glu Ile Leu Ala Leu Gln Arg Asp Leu
                85                  90                  95
Val Lys Gln Phe Gln Asn Val Gln Gln Val Glu Tyr Ser Ser Ile Arg
            100                 105                 110
Gly Phe Leu Ser Lys His Ser Ser Asp Gly Leu Arg Gln Leu Leu His
            115                 120                 125
Asn Arg Ile Thr Val Phe Leu Ser Thr Trp Asn Lys Leu Arg Arg Ser
    130                 135                 140
Leu Glu Thr Asn Gly Glu Ile Asn Leu Pro Lys Asp Tyr Cys Ser Thr
145                 150                 155                 160
Asp Leu Asp Leu Asp Thr Glu Phe Glu Ile Leu Leu Pro Arg Arg Arg
            165                 170                 175
Gly Leu Gly Leu Cys Ala Thr Ala Leu Val Ser Tyr Leu Ile Arg Leu
            180                 185                 190
His Asn Glu Ile Val Tyr Ala Val Glu Lys Leu Ser Lys Glu Asn Asn
            195                 200                 205
Ser Tyr Ser Val Asp Ala Ala Glu Val Thr Glu Leu His Val Ile Ser
    210                 215                 220
Tyr Glu Val Glu Arg Asp Leu Thr Pro Leu Ile Leu Ser Asn Cys Gln
225                 230                 235                 240
Tyr Gln Val Glu Glu Gly Arg Glu Thr Val Gln Glu Phe Asp Leu Glu
            245                 250                 255
Lys Ile Gln Arg Gln Ile Val Ser Arg Phe Leu Gln Gly Lys Pro Arg
            260                 265                 270
Leu Ser Leu Lys Gly Ile Pro Thr Leu Val Tyr Arg His Asp Trp Asn
    275                 280                 285
Tyr Glu His Leu Phe Met Asp Ile Lys Asn Lys Met Ala Gln Asp Ser
    290                 295                 300
Leu Pro Ser Ser Val Ile Ser Ala Ile Ser Gly Gln Leu Gln Ser Tyr
305                 310                 315                 320
Ser Asp Ala Cys Glu Val Leu Ser Val Val Glu Val Thr Leu Gly Phe
            325                 330                 335
Leu Ser Thr Ala Gly Gly Asp Pro Asn Met Gln Leu Asn Val Tyr Thr
            340                 345                 350
Gln Asp Ile Leu Gln Met Gly Asp Gln Thr Ile His Val Leu Lys Ala
            355                 360                 365
Leu Asn Arg Cys Gln Leu Lys His Thr Ile Ala Leu Trp Gln Phe Leu
    370                 375                 380
Ser Ala His Lys Ser Glu Gln Leu Leu Arg Leu His Lys Glu Pro Phe
385                 390                 395                 400
Gly Glu Ile Ser Ser Arg Tyr Lys Ala Asp Leu Ser Pro Glu Asn Ala
            405                 410                 415
Lys Leu Leu Ser Thr Phe Leu Asn Gln Thr Gly Leu Asp Ala Phe Leu
            420                 425                 430
Leu Glu Leu His Glu Met Ile Ile Leu Lys Leu Lys Asn Pro Gln Thr
            435                 440                 445
Gln Thr Glu Glu Arg Phe Arg Pro Gln Trp Ser Leu Arg Asp Thr Leu
    450                 455                 460
Val Ser Tyr Met Gln Thr Lys Glu Ser Glu Ile Leu Pro Glu Met Ala
465                 470                 475                 480
Ser Gln Phe Pro Glu Glu Ile Leu Leu Ala Ser Cys Val Ser Val Trp
            485                 490                 495
Lys Thr Ala Ala Val Leu Lys Trp Asn Arg Glu Met Arg
            500                 505
```

<210> 57
<211> 1760
<212> DNA
<213> Homo sapiens

<400> 57
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gctgaccgcg ctagctgcgg cttctacgct ccggcactct gagttcatca gcaaacgccc 180
tggcgtctgt cctcaccatg cctagccttt gggaccgctt ctcgtcgtcg tccacctcct 240
cttcgccctc gtccttgccc cgaactccca ccccagatcg gccgccgcgc tcagcctggg 300
ggtcggcgac ccgggaggag gggtttgacc gctccacgag cctggagagc tcggactgcg 360
agtccctgga cagcagcaac agtggcttcg ggccggagga agacacggct tacctggatg 420
gggtgtcgtt gcccgacttc gagctgctca gtgaccctga ggatgaacac ttgtgtgcca 480
acctgatgca gctgctgcag gagagcctgg cccaggcgcg gctgggctct cgacgccctg 540
cgcgcctgct gatgcctagc cagttggtaa gccaggtggg caaagaacta ctgcgcctgg 600
cctacagcga gccgtgcggc ctgcgggggg cgctgctgga cgtctgcgtg gagcagggca 660
agagctgcca cagcgtgggc cagctggcac tcgaccccag cctggtgccc accttccagc 720
tgaccctcgt gctgcgcctg gactcacgac tctggcccaa gatccagggg ctgtttagct 780
ccgccaactc tcccttcctc cctggcttca gccagtccct gacgctgagc actggcttcc 840
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ttgtttttct gatcggagca tcactactga cctgttgtag gcagctatct tacagacgca 1440
tgaatgtaag agtaggaagg ggtgggtgtc agggatcact tgggatcttt gacacttgaa 1500
aaattacacc tggcagctgc gtttaagcct tcccccatcg tgtactgcag agttgagctg 1560
gcaggggagg ggctgagagg gtgggggctg gaacccctcc cgggaggag tgccatctgg 1620
gtcttccatc tagaactgtt tacatgaaga taagatactc actgttcatg aatacacttg 1680
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aacaaaaaaa aaaaaaaaaa 1760

<210> 58
<211> 232
<212> PRT
<213> Homo sapiens

<400> 58
Met Pro Ser Leu Trp Asp Arg Phe Ser Ser Ser Ser Thr Ser Ser Ser
1   5   10   15
Pro Ser Ser Leu Pro Arg Thr Pro Thr Pro Asp Arg Pro Pro Arg Ser
            20   25   30
Ala Trp Gly Ser Ala Thr Arg Glu Glu Gly Phe Asp Arg Ser Thr Ser
        35   40   45
Leu Glu Ser Ser Asp Cys Glu Ser Leu Asp Ser Ser Asn Ser Gly Phe
    50   55   60
Gly Pro Glu Glu Asp Thr Ala Tyr Leu Asp Gly Val Ser Leu Pro Asp
65   70   75   80
Phe Glu Leu Leu Ser Asp Pro Glu Asp Glu His Leu Cys Ala Asn Leu
                85   90   95
Met Gln Leu Leu Gln Glu Ser Leu Ala Gln Ala Arg Leu Gly Ser Arg
            100   105   110

```
Arg Pro Ala Arg Leu Leu Met Pro Ser Gln Leu Val Ser Gln Val Gly
        115                 120                 125
Lys Glu Leu Leu Arg Leu Ala Tyr Ser Glu Pro Cys Gly Leu Arg Gly
    130                 135                 140
Ala Leu Leu Asp Val Cys Val Glu Gln Gly Lys Ser Cys His Ser Val
145                 150                 155                 160
Gly Gln Leu Ala Leu Asp Pro Ser Leu Val Pro Thr Phe Gln Leu Thr
                165                 170                 175
Leu Val Leu Arg Leu Asp Ser Arg Leu Trp Pro Lys Ile Gln Gly Leu
            180                 185                 190
Phe Ser Ser Ala Asn Ser Pro Phe Leu Pro Gly Phe Ser Gln Ser Leu
        195                 200                 205
Thr Leu Ser Thr Gly Phe Arg Val Ile Lys Lys Lys Leu Tyr Ser Ser
    210                 215                 220
Glu Gln Leu Leu Ile Glu Glu Cys
225                 230
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```
<210> 59
<211> 2012
<212> DNA
<213> Homo sapiens

<400> 59
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agccagtgtt ctgaaagtct ccctgttcgc acactaagca ttgcaccagg tcaatgtcga 180
ccaccgaggg ttttgggtag accaaagcac aaagaagtcc acttagagtg ggatgttcct 240
gcatcggaaa gtggctgtga ggtctcagag tacagcgtgg agatgacgga gcccgaagac 300
gtagcctcgg aagtgtacca tggcccagag ctggagtgca ccgtcggcaa cctgcttcct 360
ggaaccgtgt atcgcttccg ggtgagggct ctgaatgatg agggtatggt ccctattct 420
gatgtctcag aaattaccac tgctgcaggg cctcctggac aatgcaaagc accttgtatt 480
tcttgtacac ctgatggatg tgtcttagtg ggttgggaga gtcctgatag ttctggtgct 540
gacatctcag agtacaggtt ggaatgggga gaagatgaag aatccttaga actcatttat 600
catgggacag acacccgttt tgaaataaga gacctgttgc ctgctgcaca gtattgctgt 660
agactacagg ccttcaatca agcaggggca gggccgtaca gtgaacttgt cctttgccag 720
acgccagcgt ctgcccctga ccccgtctcc actctctgtg tcctggagga ggagcccctt 780
gatgcctacc ctgattcacc ttctgcgtgc cttgtactga actgtgaaga gccgtgcaat 840
aacggatctg aaatccttgc ttacaccatt gatctaggag acactagcat taccgtgggc 900
aacaccacca tgcatgttat gaaagatctc cttccagaaa ccacctaccg gatcagaatt 960
caggctataa atgaaattgg agctggacca tttagtcagt tcattaaagc aaaaactcgg 1020
ccattaccac ccttgcctcc taggctagaa tgtgctgctg ctggtcctca gagcctgaag 1080
ctaaaatggg gagacagtaa ctccaagaca catgctgctg aggacattgt gtacacacta 1140
cagctggagg acagaaacaa gaggtttatt tcaatctaca gaggacccag ccacacctac 1200
aaggtccaga gactgacgga attcacatgc tactccttca gaatccaggc agcaagcgag 1260
gctggagaag ggcccttctc agaaacctat accttcagca caaccaaaag tgtcccccc 1320
accatcaaag cacctcgagt aacacagtta gaaggaaatt catgtgaaat tttatgggag 1380
acggtaccat caatgaaagg tgaccctgtt aactacattc tgcaggtatt ggttggaaga 1440
gaatctgagt acaaacaggt gtacaaggga gaagaagcca cattccaaat ctcaggcctc 1500
cagaccaaca cagactacag gttccgcgta tgtgcgtgtc gtcgctgttt agacacctct 1560
caggagctaa gcggagcctt cagcccctct gcggcttttg tattacaacg aagtgaggtc 1620
atgcttacag gggacatggg gagcttagat gatcccaaaa tgaagagcat gatgcctact 1680
gatgaacagt ttgcagccat cattgtgctt ggctttgcaa ctttgtccat tttatttgcc 1740
tttatattac agtacttctt aatgaagtaa acccaacaaa actagaggta tgaattaatg 1800
ctacacattt taatacacac atttattcag atactcccct ttttaaagcc cttttgtttt 1860
ttgatttata tactctgttt tacagattta gctagaaaaa aaatgtcagt gttttggtgc 1920
acctttttga aatgcaaaac taggaaaagg ttaaactgga tttttttttt taaaaaaaaa 1980
aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aa 2012
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<210> 60
<211> 495
<212> PRT
<213> Homo sapiens

<400> 60
Met Thr Glu Pro Glu Asp Val Ala Ser Glu Val Tyr His Gly Pro Glu
1   5   10   15
Leu Glu Cys Thr Val Gly Asn Leu Leu Pro Gly Thr Val Tyr Arg Phe
            20   25   30
Arg Val Arg Ala Leu Asn Asp Gly Gly Tyr Gly Pro Tyr Ser Asp Val
        35   40   45
Ser Glu Ile Thr Thr Ala Ala Gly Pro Pro Gly Gln Cys Lys Ala Pro
    50   55   60
Cys Ile Ser Cys Thr Pro Asp Gly Cys Val Leu Val Gly Trp Glu Ser
65   70   75   80
Pro Asp Ser Ser Gly Ala Asp Ile Ser Glu Tyr Arg Leu Glu Trp Gly
                85   90   95
Glu Asp Glu Glu Ser Leu Glu Leu Ile Tyr His Gly Thr Asp Thr Arg
            100   105   110
Phe Glu Ile Arg Asp Leu Leu Pro Ala Ala Gln Tyr Cys Cys Arg Leu
        115   120   125
Gln Ala Phe Asn Gln Ala Gly Ala Gly Pro Tyr Ser Glu Leu Val Leu
    130   135   140
Cys Gln Thr Pro Ala Ser Ala Pro Asp Pro Val Ser Thr Leu Cys Val
145   150   155   160
Leu Glu Glu Glu Pro Leu Asp Ala Tyr Pro Asp Ser Pro Ser Ala Cys
                165   170   175
Leu Val Leu Asn Trp Glu Glu Pro Cys Asn Asn Gly Ser Glu Ile Leu
            180   185   ·190
Ala Tyr Thr Ile Asp Leu Gly Asp Thr Ser Ile Thr Val Gly Asn Thr
        195   200   205
Thr Met His Val Met Lys Asp Leu Leu Pro Glu Thr Thr Tyr Arg Ile
    210   215   220
Arg Ile Gln Ala Ile Asn Glu Ile Gly Ala Gly Pro Phe Ser Gln Phe
225   230   235   240
Ile Lys Ala Lys Thr Arg Pro Leu Pro Pro Leu Pro Pro Arg Leu Glu
                245   250   255
Cys Ala Ala Ala Gly Pro Gln Ser Leu Lys Leu Lys Trp Gly Asp Ser
                260   265   270
Asn Ser Lys Thr His Ala Ala Glu Asp Ile Val Tyr Thr Leu Gln Leu
            275   280   285
Glu Asp Arg Asn Lys Arg Phe Ile Ser Ile Tyr Arg Gly Pro Ser His
        290   295   300
Thr Tyr Lys Val Gln Arg Leu Thr Glu Phe Thr Cys Tyr Ser Phe Arg
305   310   315   320
Ile Gln Ala Ala Ser Glu Ala Gly Glu Gly Pro Phe Ser Glu Thr Tyr
                325   330   335
Thr Phe Ser Thr Thr Lys Ser Val Pro Pro Thr Ile Lys Ala Pro Arg
            340   345   350
Val Thr Gln Leu Glu Gly Asn Ser Cys Glu Ile Leu Trp Glu Thr Val
        355   360   365
Pro Ser Met Lys Gly Asp Pro Val Asn Tyr Ile Leu Gln Val Leu Val
    370   375   380
Gly Arg Glu Ser Glu Tyr Lys Gln Val Tyr Lys Gly Glu Glu Ala Thr
385   390   395   400
Phe Gln Ile Ser Gly Leu Gln Thr Asn Thr Asp Tyr Arg Phe Arg Val
            405   , 410   415
Cys Ala Cys Arg Arg Cys Leu Asp Thr Ser Gln Glu Leu Ser Gly Ala

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          420                    425                    430
Phe Ser Pro Ser Ala Ala Phe Val Leu Gln Arg Ser Glu Val Met Leu
        435                    440                    445
Thr Gly Asp Met Gly Ser Leu Asp Asp Pro Lys Met Lys Ser Met Met
    450                    455                    460
Pro Thr Asp Glu Gln Phe Ala Ala Ile Ile Val Leu Gly Phe Ala Thr
465                    470                    475                    480
Leu Ser Ile Leu Phe Ala Phe Ile Leu Gln Tyr Phe Leu Met Lys
            485                    490                    495
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<210> 61
<211> 2384
<212> DNA
<213> Homo sapiens

<400> 61

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atcgccctaa aggactggca ttcactgatg tggatgtcga ttccatcaaa attgcttggg 180
aaagcccaca ggggcaagtt tccaggtaca gggtgaccta ctcgagccct gaggatggaa 240
tccatgagct attccctgca cctgatggtg aagaagacac tgcagagctg caaggcctca 300
gaccgggttc tgagtacaca gtcagtgtgg ttgccttgca cgatgatatg gagagccagc 360
ccctgattgg aacccagtcc acagctattc ctgcaccaac tgacctgaag ttcactcagg 420
tcacacccac aagcctgagc gcccagtgga caccacccaa tgttcagctc actggatatc 480
gagtgcgggt gacccccaag gagaagaccg gaccaatgaa agaaatcaac cttgctcctg 540
acagctcatc cgtggttgta tcaggactta tggtggccac caaatatgaa gtgagtgtct 600
atgctcttaa ggacactttg acaagcagac cagctcaggg tgttgtcacc actctggaga 660
atgtcagccc accaagaagg gctcgtgtga cagatgctac tgagaccacc atcaccatta 720
gctggagaac caagactgag acgatcactg gcttccaagt tgatgccgtt ccagccaatg 780
gccagactcc aatccagaga accatcaagc cagatgtcag aagctacacc atcacaggtt 840
tacaaccagg cactgactac aagatctacc tgtacacctt gaatgacaat gctcggagct 900
cccctgtggt catcgacgcc tccactgcca ttgatgcacc atccaacctg cgtttcctgg 960
ccaccacacc caattccttg ctggtatcat ggcagccgcc acgtgccagg attaccggct 1020
acatcatcaa gtatgagaag cctgggtctc ctcccagaga agtggtccct cggccccgcc 1080
ctggtgtcac agaggctact attactggcc tggaaccggg aaccgaatat acaatttatg 1140
tcattgccct gaagaataat cagaagagcg agcccctgat tggaaggaaa aagacagacg 1200
agcttcccca actggtaacc cttccacacc ccaatcttca tggaccagag atcttggatg 1260
ttccttccac agttcaaaag acccctttcg tcacccaccc tgggtatgac actggaaatg 1320
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aggaacatgg ttttaggcgg accacaccgc ccacaacggc cacccccata aggcataggc 1440
caagaccata cccgccgaat gtaggtgagg aaatccaaat tggtcacatt cccagggaag 1500
atgtagacta tcacctgtac ccacacggtc cggggctcaa tccaaatgcc tctacaggac 1560
aagaagctct ctctcagaca accatctcat gggccccatt ccaggacact tctgagtaca 1620
tcatttcatg tcatcctgtt ggcactgatg aagaaccctt acagttcagg gttcctggaa 1680
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cccattatgc cgttggagat gagtgggaac gaatgtctga atcaggcttt aaactgttgt 1920
gccagtgctt aggctttgga agtggtcatt tcagatgtga ttcatctaga tggtgccatg 1980
acaatggtgt gaactacaag attggagaga agtgggaccg tcaggagaa aatggccaga 2040
tgatgagctg cacatgtctt gggaacggaa aaggagaatt caagtgtgac cctcatgagg 2100
caacgtgtta cgatgatggg aagacatacc acgtaggaga acagtggcag aaggaatatc 2160
tcggtgccat ttgctcctgc acatgctttg gaggccagcg gggctggcgc tgtgacaact 2220
gccgcagacc tgggggtgaa cccagtcccg aaggcactac tggccagtcc tacaaccagt 2280
attctcagag ataccatcag agaacaaaca ctaatgttaa ttgcccaatt gagtgcttca 2340
tgcctttaga tgtacaggct gacagagaag attcccgaga gtaa                2384
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<210> 62

<211> 793
<212> PRT
<213> Homo sapiens

<400> 62
Gln Thr Glu Met Thr Ile Glu Gly Leu Gln Pro Thr Val Glu Tyr Val
1   5   10   15
Val Ser Val Tyr Ala Gln Asn Pro Ser Gly Glu Ser Gln Pro Leu Val
            20   25   30
Gln Thr Ala Val Thr Asn Ile Asp Arg Pro Lys Gly Leu Ala Phe Thr
        35   40   45
Asp Val Asp Val Asp Ser Ile Lys Ile Ala Trp Glu Ser Pro Gln Gly
    50   55   60
Gln Val Ser Arg Tyr Arg Val Thr Tyr Ser Ser Pro Glu Asp Gly Ile
65   70   75   80
His Glu Leu Phe Pro Ala Pro Asp Gly Glu Glu Asp Thr Ala Glu Leu
                85   90   95
Gln Gly Leu Arg Pro Gly Ser Glu Tyr Thr Val Ser Val Val Ala Leu
            100   105   110
His Asp Asp Met Glu Ser Gln Pro Leu Ile Gly Thr Gln Ser Thr Ala
        115   120   125
Ile Pro Ala Pro Thr Asp Leu Lys Phe Thr Gln Val Thr Pro Thr Ser
    130   135   140
Leu Ser Ala Gln Trp Thr Pro Pro Asn Val Gln Leu Thr Gly Tyr Arg
145   150   155   160
Val Arg Val Thr Pro Lys Glu Lys Thr Gly Pro Met Lys Glu Ile Asn
                165   170   175
Leu Ala Pro Asp Ser Ser Ser Val Val Val Ser Gly Leu Met Val Ala
            180   185   190
Thr Lys Tyr Glu Val Ser Val Tyr Ala Leu Lys Asp Thr Leu Thr Ser
            195   200   205
Arg Pro Ala Gln Gly Val Val Thr Thr Leu Glu Asn Val Ser Pro Pro
    210   215   220
Arg Arg Ala Arg Val Thr Asp Ala Thr Glu Thr Thr Ile Thr Ile Ser
225   230   235   240
Trp Arg Thr Lys Thr Glu Thr Ile Thr Gly Phe Gln Val Asp Ala Val
                245   250   255
Pro Ala Asn Gly Gln Thr Pro Ile Gln Arg Thr Ile Lys Pro Asp Val
            260   265   270
Arg Ser Tyr Thr Ile Thr Gly Leu Gln Pro Gly Thr Asp Tyr Lys Ile
        275   280   285
Tyr Leu Tyr Thr Leu Asn Asp Asn Ala Arg Ser Ser Pro Val Val Ile
    290   295   300
Asp Ala Ser Thr Ala Ile Asp Ala Pro Ser Asn Leu Arg Phe Leu Ala
305   310   315   320
Thr Thr Pro Asn Ser Leu Leu Val Ser Trp Gln Pro Pro Arg Ala Arg
                325   330   335
Ile Thr Gly Tyr Ile Ile Lys Tyr Glu Lys Pro Gly Ser Pro Pro Arg
            340   345   350
Glu Val Val Pro Arg Pro Arg Pro Gly Val Thr Glu Ala Thr Ile Thr
        355   360   365
Gly Leu Glu Pro Gly Thr Glu Tyr Thr Ile Tyr Val Ile Ala Leu Lys
    370   375   380
Asn Asn Gln Lys Ser Glu Pro Leu Ile Gly Arg Lys Lys Thr Asp Glu
385   390   395   400
Leu Pro Gln Leu Val Thr Leu Pro His Pro Asn Leu His Gly Pro Glu
                405   410   415
Ile Leu Asp Val Pro Ser Thr Val Gln Lys Thr Pro Phe Val Thr His
            420   425   430

```
Pro Gly Tyr Asp Thr Gly Asn Gly Ile Gln Leu Pro Gly Thr Ser Gly
        435             440             445
Gln Gln Pro Ser Val Gly Gln Met Ile Phe Glu Glu His Gly Phe
    450             455             460
Arg Arg Thr Thr Pro Pro Thr Thr Ala Thr Pro Ile Arg His Arg Pro
465             470             475             480
Arg Pro Tyr Pro Pro Asn Val Gly Glu Glu Ile Gln Ile Gly His Ile
            485             490             495
Pro Arg Glu Asp Val Asp Tyr His Leu Tyr Pro His Gly Pro Gly Leu
            500             505             510
Asn Pro Asn Ala Ser Thr Gly Gln Glu Ala Leu Ser Gln Thr Thr Ile
            515             520             525
Ser Trp Ala Pro Phe Gln Asp Thr Ser Glu Tyr Ile Ile Ser Cys His
    530             535             540
Pro Val Gly Thr Asp Glu Glu Pro Leu Gln Phe Arg Val Pro Gly Thr
545             550             555             560
Ser Thr Ser Ala Thr Leu Thr Gly Leu Thr Arg Gly Ala Thr Tyr Asn
            565             570             575
Ile Ile Val Glu Ala Leu Lys Asp Gln Gln Arg His Lys Val Arg Glu
            580             585             590
Glu Val Val Thr Val Gly Asn Ser Val Asn Glu Gly Leu Asn Gln Pro
    595             600             605
Thr Asp Asp Ser Cys Phe Asp Pro Tyr Thr Val Ser His Tyr Ala Val
    610             615             620
Gly Asp Glu Trp Glu Arg Met Ser Glu Ser Gly Phe Lys Leu Leu Cys
625             630             635             640
Gln Cys Leu Gly Phe Gly Ser Gly His Phe Arg Cys Asp Ser Ser Arg
            645             650             655
Trp Cys His Asp Asn Gly Val Asn Tyr Lys Ile Gly Glu Lys Trp Asp
            660             665             670
Arg Gln Gly Glu Asn Gly Gln Met Met Ser Cys Thr Cys Leu Gly Asn
    675             680             685
Gly Lys Gly Glu Phe Lys Cys Asp Pro His Glu Ala Thr Cys Tyr Asp
    690             695             700
Asp Gly Lys Thr Tyr His Val Gly Glu Gln Trp Gln Lys Glu Tyr Leu
705             710             715             720
Gly Ala Ile Cys Ser Cys Thr Cys Phe Gly Gly Gln Arg Gly Trp Arg
            725             730             735
Cys Asp Asn Cys Arg Arg Pro Gly Gly Glu Pro Ser Pro Glu Gly Thr
            740             745             750
Thr Gly Gln Ser Tyr Asn Gln Tyr Ser Gln Arg Tyr His Gln Arg Thr
    755             760             765
Asn Thr Asn Val Asn Cys Pro Ile Glu Cys Phe Met Pro Leu Asp Val
770             775             780
Gln Ala Asp Arg Glu Asp Ser Arg Glu
785             790
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<210> 63
<211> 7680
<212> DNA
<213> Homo sapiens

<400> 63
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gacctaccta ggtaatgtgt tggtttgtac ttgttatgga ggaagccgag gttttaactg   180
cgaaagtaaa cctgaagctg aagagacttg ctttgacaag tacactggga acacttaccg   240
agtgggtgac acttatgagc gtcctaaaga ctccatgatc tgggactgta cctgcatcgg   300
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ggctgggcga gggagaataa gctgtaccat cgcaaaccgc tgccatgaag ggggtcagtc 360
ctacaagatt ggtgacacct ggaggagacc acatgagact ggtggttaca tgttagagtg 420
tgtgtgtctt ggtaatggaa aaggagaatg gacctgcaag cccatagctg agaagtgttt 480
tgatcatgct gctgggactt cctatgtggt cggagaaacg tgggagaagc cctaccaagg 540
ctggatgatg gtagattgta cttgcctggg agaaggcagc ggacgcatca cttgcacttc 600
tagaaataga tgcaacgatc aggacacaag gacatcctat agaattggag acacctggag 660
caagaaggat aatcgaggaa acctgctcca gtgcatctgc acaggcaacg gccgaggaga 720
gtggaagtgt gagaggcaca cctctgtgca gaccacatcg agcggatctg gcccttcac 780
cgatgttcgt gcagctgttt accaaccgca gcctcacccc cagcctcctc cctatggcca 840
ctgtgtcaca gacagtggtg tggtctactc tgtggggatg cagtggttga agacacaagg 900
aaataagcaa atgctttgca cgtgcctggg caacggagtc agctgccaag agacagctgt 960
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caaccacaat tacactgatt gcacttctga gggcagaaga gacaacatga agtggtgtgg 1260
gaccacacag aactatgatg ccgaccagaa gtttgggttc tgccccatgg ctgcccacga 1320
ggaaatctgc acaaccaatg aagggtcat gtaccgcatt ggagatcagt gggataagca 1380
gcatgacatg ggtcacatga tgaggtgcac gtgtgttggg aatggtcgtg gggaatggac 1440
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gaacgacaca ttccacaagc gtcatgaaga ggggcacatg ctgaactgta catgcttcgg 1560
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catccagtgg aatgcaccac agccatctca catttccaag tacattctca ggtggagacc 1860
taaaaattct gtaggccgtt ggaaggaagc taccatacca ggccacttaa actcctacac 1920
catcaaaggc ctgaagcctg gtgtggtata cgagggccag ctcatcagca tccagcagta 1980
cggccaccaa gaagtgactc gctttgactt caccaccacc agcaccagca cacctgtgac 2040
cagcaacacc gtgacaggag agacgactcc ctttttctcct cttgtggcca cttctgaatc 2100
tgtgaccgaa atcacagcca gtagctttgt ggtctcctgg gtctcagctt ccgacaccgt 2160
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tcttccaagc acagccactt ctgtgaacat ccctgacctg cttcctggcc gaaaatacat 2280
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<211> 2328
<212> PRT
<213> Homo sapiens
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Gln Ile Asn Gln Gln Trp Glu Arg Thr Tyr Leu Gly Asn Val Leu Val
            35                  40                  45
Cys Thr Cys Tyr Gly Gly Ser Arg Gly Phe Asn Cys Glu Ser Lys Pro
        50                  55                  60
Glu Ala Glu Glu Thr Cys Phe Asp Lys Tyr Thr Gly Asn Thr Tyr Arg
65                  70                  75                  80
Val Gly Asp Thr Tyr Glu Arg Pro Lys Asp Ser Met Ile Trp Asp Cys
                85                  90                  95
Thr Cys Ile Gly Ala Gly Arg Gly Arg Ile Ser Cys Thr Ile Ala Asn
            100                 105                 110
Arg Cys His Glu Gly Gly Gln Ser Tyr Lys Ile Gly Asp Thr Trp Arg
        115                 120                 125
Arg Pro His Glu Thr Gly Gly Tyr Met Leu Glu Cys Val Cys Leu Gly
    130                 135                 140
Asn Gly Lys Gly Glu Trp Thr Cys Lys Pro Ile Ala Glu Lys Cys Phe
145                 150                 155                 160
Asp His Ala Ala Gly Thr Ser Tyr Val Val Gly Glu Thr Trp Glu Lys
    .           165                 170                 175
Pro Tyr Gln Gly Trp Met Met Val Asp Cys Thr Cys Leu Gly Glu Gly
            180                 185                 190
Ser Gly Arg Ile Thr Cys Thr Ser Arg Asn Arg Cys Asn Asp Gln Asp
        195                 200                 205
Thr Arg Thr Ser Tyr Arg Ile Gly Asp Thr Trp Ser Lys Lys Asp Asn
    210                 215                 220
Arg Gly Asn Leu Leu Gln Cys Ile Cys Thr Gly Asn Gly Arg Gly Glu
225             .   230                 235             .       240
Trp Lys Cys Glu Arg His Thr Ser Val Gln Thr Thr Ser Ser Gly Ser
            245                 250                 255
Gly Pro Phe Thr Asp Val Arg Ala Ala Val Tyr Gln Pro Gln Pro His
            260                 265                 270
Pro Gln Pro Pro Pro Tyr Gly His Cys Val Thr Asp Ser Gly Val Val
        275                 280                 285
Tyr Ser Val Gly Met Gln Trp Leu Lys Thr Gln Gly Asn Lys Gln Met
    290                 295                 300
Leu Cys Thr Cys Leu Gly Asn Gly Val Ser Cys Gln Glu Thr Ala Val
305                 310                 315                 320
Thr Gln Thr Tyr Gly Gly Asn Leu Asn Gly Glu Pro Cys Val Leu Pro
            325                 330                 335
Phe Thr Tyr Asn Gly Arg Thr Phe Tyr Ser Cys Thr Thr Glu Gly Arg
            340                 345                 350
Gln Asp Gly His Leu Trp Cys Ser Thr Thr Ser Asn Tyr Glu Gln Asp
        355             .       360                 365
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Gln Lys Tyr Ser Phe Cys Thr Asp His Thr Val Leu Val Gln Thr Gln
    370             375             380
Gly Gly Asn Ser Asn Gly Ala Leu Cys His Phe Pro Phe Leu Tyr Asn
385             390             395             400
Asn His Asn Tyr Thr Asp Cys Thr Ser Glu Gly Arg Arg Asp Asn Met
            405             410             415
Lys Trp Cys Gly Thr Thr Gln Asn Tyr Asp Ala Asp Gln Lys Phe Gly
        420             425             430
Phe Cys Pro Met Ala Ala His Glu Glu Ile Cys Thr Thr Asn Glu Gly
        435             440             445
Val Met Tyr Arg Ile Gly Asp Gln Trp Asp Lys Gln His Asp Met Gly
    450             455             460
His Met Met Arg Cys Thr Cys Val Gly Asn Gly Arg Gly Glu Trp Thr
465             470             475             480
Cys Ile Ala Tyr Ser Gln Leu Arg Asp Gln Cys Ile Val Asp Asp Ile
            485             490             495
Thr Tyr Asn Val Asn Asp Thr Phe His Lys Arg His Glu Glu Gly His
            500             505             510
Met Leu Asn Cys Thr Cys Phe Gly Gln Gly Arg Gly Arg Trp Lys Cys
        515             520             525
Asp Pro Val Asp Gln Cys Gln Asp Ser Glu Thr Gly Thr Phe Tyr Gln
    530             535             540
Ile Gly Asp Ser Trp Glu Lys Tyr Val His Gly Val Arg Tyr Gln Cys
545             550             555             560
Tyr Cys Tyr Gly Arg Gly Ile Gly Glu Trp His Cys Gln Pro Leu Gln
            565             570             575
Thr Tyr Pro Ser Ser Ser Gly Pro Val Glu Val Phe Ile Thr Glu Thr
        580             585             590
Pro Ser Gln Pro Asn Ser His Pro Ile Gln Trp Asn Ala Pro Gln Pro
    595             600             605
Ser His Ile Ser Lys Tyr Ile Leu Arg Trp Arg Pro Lys Asn Ser Val
610             615             620
Gly Arg Trp Lys Glu Ala Thr Ile Pro Gly His Leu Asn Ser Tyr Thr
625             630             635             640
Ile Lys Gly Leu Lys Pro Gly Val Val Tyr Glu Gly Gln Leu Ile Ser
            645             650             655
Ile Gln Gln Tyr Gly His Gln Glu Val Thr Arg Phe Asp Phe Thr Thr
        660             665             670
Thr Ser Thr Ser Thr Pro Val Thr Ser Asn Thr Val Thr Gly Glu Thr
    675             680             685
Thr Pro Phe Ser Pro Leu Val Ala Thr Ser Glu Ser Val Thr Glu Ile
    690             695             700
Thr Ala Ser Ser Phe Val Val Ser Trp Val Ser Ala Ser Asp Thr Val
705             710             715             720
Ser Gly Phe Arg Val Glu Tyr Glu Leu Ser Glu Glu Gly Asp Glu Pro
            725             730             735
Gln Tyr Leu Asp Leu Pro Ser Thr Ala Thr Ser Val Asn Ile Pro Asp
        740             745             750
Leu Leu Pro Gly Arg Lys Tyr Ile Val Asn Val Tyr Gln Ile Ser Glu
        755             760             765
Asp Gly Glu Gln Ser Leu Ile Leu Ser Thr Ser Gln Thr Thr Ala Pro
    770             775             780
Asp Ala Pro Pro Asp Pro Thr Val Asp Gln Val Asp Asp Thr Ser Ile
785             790             795             800
Val Val Arg Trp Ser Arg Pro Gln Ala Pro Ile Thr Gly Tyr Arg Ile
            805             810             815
Val Tyr Ser Pro Ser Val Glu Gly Ser Ser Thr Glu Leu Asn Leu Pro
            820             825             830
Glu Thr Ala Asn Ser Val Thr Leu Ser Asp Leu Gln Pro Gly Val Gln
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Tyr Asn Ile Thr Ile Tyr Ala Val Glu Glu Asn Gln Glu Ser Thr Pro
    850                     855                    860
Val Val Ile Gln Gln Glu Thr Thr Gly Thr Pro Arg Ser Asp Thr Val
865                     870                    875                    880
Pro Ser Pro Arg Asp Leu Gln Phe Val Glu Val Thr Asp Val Lys Val
                885                     890                    895
Thr Ile Met Trp Thr Pro Pro Glu Ser Ala Val Thr Gly Tyr Arg Val
            900                     905                    910
Asp Val Ile Pro Val Asn Leu Pro Gly Glu His Gly Gln Arg Leu Pro
        915                     920                    925
Ile Ser Arg Asn Thr Phe Ala Glu Val Thr Gly Leu Ser Pro Gly Val
    930                     935                    940
Thr Tyr Tyr Phe Lys Val Phe Ala Val Ser His Gly Arg Glu Ser Lys
945                     950                    955                    960
Pro Leu Thr Ala Gln Gln Thr Thr Lys Leu Asp Ala Pro Thr Asn Leu
                965                     970                    975
Gln Phe Val Asn Glu Thr Asp Ser Thr Val Leu Val Arg Trp Thr Pro
            980                     985                    990
Pro Arg Ala Gln Ile Thr Gly Tyr Arg Leu Thr Val Gly Leu Thr Arg
            995                    1000                   1005
Arg Gly Gln Pro Arg Gln Tyr Asn Val Gly Pro Ser Val Ser Lys Tyr
    1010                    1015                   1020
Pro Leu Arg Asn Leu Gln Pro Ala Ser Glu Tyr Thr Val Ser Leu Val
1025                    1030                   1035                   1040
Ala Ile Lys Gly Asn Gln Glu Ser Pro Lys Ala Thr Gly Val Phe Thr
                1045                    1050                   1055
Thr Leu Gln Pro Gly Ser Ser Ile Pro Pro Tyr Asn Thr Glu Val Thr
            1060                    1065                   1070
Glu Thr Thr Ile Val Ile Thr Trp Thr Pro Ala Pro Arg Ile Gly Phe
            1075                    1080                   1085
Lys Leu Gly Val Arg Pro Ser Gln Gly Gly Glu Ala Pro Arg Glu Val
        1090                    1095                   1100
Thr Ser Asp Ser Gly Ser Ile Val Val Ser Gly Leu Thr Pro Gly Val
1105                    1110                   1115                   1120
Glu Tyr Val Tyr Thr Ile Gln Val Leu Arg Asp Gly Gln Glu Arg Asp
            1125                    1130                   1135
Ala Pro Ile Val Asn Lys Val Val Thr Pro Leu Ser Pro Pro Thr Asn
            1140                    1145                   1150
Leu His Leu Glu Ala Asn Pro Asp Thr Gly Val Leu Thr Val Ser Trp
        1155                    1160                   1165
Glu Arg Ser Thr Thr Pro Asp Ile Thr Gly Tyr Arg Ile Thr Thr Thr
1170                    1175                   1180
Pro Thr Asn Gly Gln Gln Gly Asn Ser Leu Glu Glu Val Val His Ala
1185                    1190                   1195                   1200
Asp Gln Ser Ser Cys Thr Phe Asp Asn Leu Ser Pro Gly Leu Glu Tyr
                1205                    1210                   1215
Asn Val Ser Val Tyr Thr Val Lys Asp Asp Lys Glu Ser Val Pro Ile
            1220                    1225                   1230
Ser Asp Thr Ile Ile Pro Ala Val Pro Pro Pro Thr Asp Leu Arg Phe
            1235                    1240                   1245
Thr Asn Ile Gly Pro Asp Thr Met Arg Val Thr Trp Ala Pro Pro Pro
    1250                    1255                   1260
Ser Ile Asp Leu Thr Asn Phe Leu Val Arg Tyr Ser Pro Val Lys Asn
1265                    1270                   1275                   1280
Glu Glu Asp Val Ala Glu Leu Ser Ile Ser Pro Ser Asp Asn Ala Val
                1285                    1290                   1295
Val Leu Thr Asn Leu Leu Pro Gly Thr Glu Tyr Val Val Ser Val Ser
            1300                    1305                   1310
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Ser Val Tyr Glu Gln His Glu Ser Thr Pro Leu Arg Gly Arg Gln Lys
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Thr Gly Leu Asp Ser Pro Thr Gly Ile Asp Phe Ser Asp Ile Thr Ala
        1330   1335   1340
Asn Ser Phe Thr Val His Trp Ile Ala Pro Arg Ala Thr Ile Thr Gly
1345   1350   1355   1360
Tyr Arg Ile Arg His His Pro Glu His Phe Ser Gly Arg Pro Arg Glu
                1365   1370   1375
Asp Arg Val Pro His Ser Arg Asn Ser Ile Thr Leu Thr Asn Leu Thr
            1380   1385   1390
Pro Gly Thr Glu Tyr Val Val Ser Ile Val Ala Leu Asn Gly Arg Glu
        1395   1400   1405
Glu Ser Pro Leu Leu Ile Gly Gln Gln Ser Thr Val Ser Asp Val Pro
        1410   1415   1420
Arg Asp Leu Glu Val Val Ala Ala Thr Pro Thr Ser Leu Leu Ile Ser
1425   1430   1435   1440
Trp Asp Ala Pro Ala Val Thr Val Arg Tyr Tyr Arg Ile Thr Tyr Gly
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Glu Thr Gly Gly Asn Ser Pro Val Gln Glu Phe Thr Val Pro Gly Ser
                1460   1465   .1470
Lys Ser Thr Ala Thr Ile Ser Gly Leu Lys Pro Gly Val Asp Tyr Thr
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Ile Thr Val Tyr Ala Val Thr Gly Arg Gly Asp Ser Pro Ala Ser Ser
        1490   1495   1500
Lys Pro Ile Ser Ile Asn Tyr Arg Thr Glu Ile Asp Lys Pro Ser Gln
1505   1510   1515   1520
Met Gln Val Thr Asp Val Gln Asp Asn Ser Ile Ser Val Lys Trp Leu
            1525   1530   1535
Pro Ser Ser Ser Pro Val Thr Gly Tyr Arg Val Thr Thr Thr Pro Lys
            1540   1545   1550
Asn Gly Pro Gly Pro Thr Lys Thr Lys Thr Ala Gly Pro Asp Gln Thr
            1555   1560   1565
Glu Met Thr Ile Glu Gly Leu Gln Pro Thr Val Glu Tyr Val Val Ser
        1570   1575   1580
Val Tyr Ala Gln Asn Pro Ser Gly Glu Ser Gln Pro Leu Val Gln Thr
1585   1590   1595   1600
Ala Val Thr Asn Ile Asp Arg Pro Lys Gly Leu Ala Phe Thr Asp Val
            1605   1610   1615
Asp Val Asp Ser Ile Lys Ile Ala Trp Glu Ser Pro Gln Gly Gln Val
            1620   1625   1630
Ser Arg Tyr Arg Val Thr Tyr Ser Ser Pro Glu Asp Gly Ile His Glu
        1635   1640   1645
Leu Phe Pro Ala Pro Asp Gly Glu Glu Asp Thr Ala Glu Leu Gln Gly
        1650   1655   1660
Leu Arg Pro Gly Ser Glu Tyr Thr Val Ser Val Val Ala Leu His Asp
1665   1670   1675   1680
Asp Met Glu Ser Gln Pro Leu Ile Gly Thr Gln Ser Thr Ala Ile Pro
                1685   1690   1695
Ala Pro Thr Asp Leu Lys Phe Thr Gln Val Thr Pro Thr Ser Leu Ser
            1700   1705   1710
Ala Gln Trp Thr Pro Pro Asn Val Gln Leu Thr Gly Tyr Arg Val Arg
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Val Thr Pro Lys Glu Lys Thr Gly Pro Met Lys Glu Ile Asn Leu Ala
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Pro Asp Ser Ser Ser Val Val Val Ser Gly Leu Met Val Ala Thr Lys
1745   .   1750   1755   1760
Tyr Glu Val Ser Val Tyr Ala Leu Lys Asp Thr Leu Thr Ser Arg Pro
                1765   1770   1775
Ala Gln Gly Val Val Thr Thr Leu Glu Asn Val Ser Pro Pro Arg Arg

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Thr Lys Thr Glu Thr Ile Thr Gly Phe Gln Val Asp Ala Val Pro Ala
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Asn Gly Gln Thr Pro Ile Gln Arg Thr Ile Lys Pro Asp Val Arg Ser
1825                    1830                    1835                    1840
Tyr Thr Ile Thr Gly Leu Gln Pro Gly Thr Asp Tyr Lys Ile Tyr Leu
              1845                    1850                    1855
Tyr Thr Leu Asn Asp Asn Ala Arg Ser Ser Pro Val Val Ile Asp Ala
              1860                    1865                    1870
Ser Thr Ala Ile Asp Ala Pro Ser Asn Leu Arg Phe Leu Ala Thr Thr
         1875                    1880                    1885
Pro Asn Ser Leu Leu Val Ser Trp Gln Pro Pro Arg Ala Arg Ile Thr
         1890                    1895                    1900
Gly Tyr Ile Ile Lys Tyr Glu Lys Pro Gly Ser Pro Pro Arg Glu Val
1905                    1910                    1915                    1920
Val Pro Arg Pro Arg Pro Gly Val Thr Glu Ala Thr Ile Thr Gly Leu
              1925                    1930                    1935
Glu Pro Gly Thr Glu Tyr Thr Ile Tyr Val Ile Ala Leu Lys Asn Asn
              1940                    1945                    1950
Gln Lys Ser Glu Pro Leu Ile Gly Arg Lys Lys Thr Asp Glu Leu Pro
              1955                    1960                    1965
Gln Leu Val Thr Leu Pro His Pro Asn Leu His Gly Pro Glu Ile Leu
         1970                    1975                    1980
Asp Val Pro Ser Thr Val Gln Lys Thr Pro Phe Val Thr His Pro Gly
1985                    1990                    1995                    2000
Tyr Asp Thr Gly Asn Gly Ile Gln Leu Pro Gly Thr Ser Gly Gln Gln
              2005                    2010                    2015
Pro Ser Val Gly Gln Gln Met Ile Phe Glu Glu His Gly Phe Arg Arg
              2020                    2025                    2030
Thr Thr Pro Pro Thr Thr Ala Thr Pro Ile Arg His Arg Pro Arg Pro
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         2050                    2055                    2060
Trp Ala Pro Phe Gln Asp Thr Ser Glu Tyr Ile Ile Ser Cys His Pro
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Val Gly Thr Asp Glu Glu Pro Leu Gln Phe Arg Val Pro Gly Thr Ser
              2085                    2090                    2095
Thr Ser Ala Thr Leu Thr Gly Leu Thr Arg Gly Ala Thr Tyr Asn Ile
              2100                    2105                    2110
Ile Val Glu Ala Leu Lys Asp Gln Gln Arg His Lys Val Arg Glu Glu
         2115                    2120                    2125
Val Val Thr Val Gly Asn Ser Val Asn Glu Gly Leu Asn Gln Pro Thr
         2130                    2135                    2140
Asp Asp Ser Cys Phe Asp Pro Tyr Thr Val Ser His Tyr Ala Val Gly
2145                    2150                    2155                    2160
Asp Glu Trp Glu Arg Met Ser Glu Ser Gly Phe Lys Leu Leu Cys Gln
              2165                    2170                    2175
Cys Leu Gly Phe Gly Ser Gly His Phe Arg Cys Asp Ser Ser Arg Trp
              2180                    2185                    2190
Cys His Asp Asn Gly Val Asn Tyr Lys Ile Gly Glu Lys Trp Asp Arg
              2195                    2200                    2205
Gln Gly Glu Asn Gly Gln Met Met Ser Cys Thr Cys Leu Gly Asn Gly
         2210                    2215                    2220
Lys Gly Glu Phe Lys Cys Asp Pro His Glu Ala Thr Cys Tyr Asp Asp
2225                    2230                    2235                    2240
Gly Lys Thr Tyr His Val Gly Glu Gln Trp Gln Lys Glu Tyr Leu Gly
              2245                    2250                    2255
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Asp Asn Cys Arg Arg Pro Gly Gly Glu Pro Ser Pro Glu Gly Thr Thr
        2275            2280            2285
Gly Gln Ser Tyr Asn Gln Tyr Ser Gln Arg Tyr His Gln Arg Thr Asn
    2290            2295            2300
Thr Asn Val Asn Cys Pro Ile Glu Cys Phe Met Pro Leu Asp Val Gln
2305            2310            2315            2320
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aagggacgct cggggggacgc tgttcctgag gtgtcgccgc ctccctgtcc tcgccctccg 180
cggtgggggga gaaacccagg agcgaagccc agagcccgcg gcgcggccgg cggacgaacg 240
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tgaatattaa agctcttcat ttctggaaaa aaaaaaaaaa aaaa 1844
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<210> 66
<211> 326
<212> PRT
<213> Homo sapiens

<400> 66
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 1           5               10              15
Ser Gly Ser Pro Ala His Ala Glu Ser Tyr Ser Ser Gly Gly Gly Gly
```

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                      20                    25                    30
Gln Gln Lys Phe Arg Val Asp Met Pro Gly Ser Gly Ser Ala Phe Ile
            35                    40                    45
Pro Thr Ile Asn Ala Ile Thr Thr Ser Gln Asp Leu Gln Trp Met Val
        50                    55                    60
Gln Pro Thr Val Ile Thr Ser Met Ser Asn Pro Tyr Pro Arg Ser His
65                      70                    75                    80
Pro Tyr Ser Pro Leu Pro Gly Leu Ala Ser Val Pro Gly His Met Ala
                85                    90                    95
Leu Pro Arg Pro Gly Val Ile Lys Thr Ile Gly Thr Thr Val Gly Arg
            100                   105                   110
Arg Arg Arg Asp Glu Gln Leu Ser Pro Glu Glu Glu Glu Lys Arg Arg
            115                   120                   125
Ile Arg Arg Glu Arg Asn Lys Leu Ala Ala Ala Lys Cys Arg Asn Arg
        130                   135                   140
Arg Arg Glu Leu Thr Glu Lys Leu Gln Ala Glu Thr Glu Glu Leu Glu
145                   150                   155                   160
Glu Glu Lys Ser Gly Leu Gln Lys Glu Ile Ala Glu Leu Gln Lys Glu
                165                   170                   175
Lys Glu Lys Leu Glu Phe Met Leu Val Ala His Gly Pro Val Cys Lys
            180                   185                   190
Ile Ser Pro Glu Glu Arg Arg Ser Pro Pro Ala Pro Gly Leu Gln Pro
            195                   200                   205
Met Arg Ser Gly Gly Gly Ser Val Gly Ala Val Val Val Lys Gln Glu
        210                   215                   220
Pro Leu Glu Glu Asp Ser Pro Ser Ser Ser Ser Ala Gly Leu Asp Lys
225                   230                   235                   240
Ala Gln Arg Ser Val Ile Lys Pro Ile Ser Ile Ala Gly Gly Phe Tyr
                245                   250                   255
Gly Glu Glu Pro Leu His Thr Pro Ile Val Val Thr Ser Thr Pro Ala
            260                   265                   270
Val Thr Pro Gly Thr Ser Asn Leu Val Phe Thr Tyr Pro Ser Val Leu
        275                   280                   285
Glu Gln Glu Ser Pro Ala Ser Pro Ser Glu Ser Cys Ser Lys Ala His
        290                   295                   300
Arg Arg Ser Ser Ser Ser Gly Asp Gln Ser Ser Asp Ser Leu Asn Ser
305                   310                   315                   320
Pro Thr Leu Leu Ala Leu
            325
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<210> 67
<211> 3602
<212> DNA
<213> Homo sapiens

<220>
<221> misc_feature
<222> 2087, 2093, 2098
<223> n = A,T,C or G

<400> 67
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aagggacgct cgggggacgc tgttcctgag gtgtcgccgc ctccctgtcc tcgccctccg 180
cggtggggga gaaacccagg agcgaagccc agagcccgcg gcgcggccgg cggacgaacg 240
agcgcgcagc agccggtgcg cggccgcggc gagggcgggg gaagaaaaac accctgtttc 300
ctctccggcc cccaccgcgg atcatgtacc aggattatcc cgggaacttt gacacctcgt 360
cccgggggcag cagcggctct cctgcgcacg ccgagtccta ctccagcggc ggcggcggcc 420
```

```
agcagaaatt ccgggtagat atgcctggct caggcagtgc attcatcccc accatcaacg 480
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ccaacccata ccctcgctcg caccectaca gccccctgcc gggcctggcc tctgtccctg 600
gacacatggc cctcccaaga cctggcgtga tcaagaccat tggcaccacc gtgggccgca 660
ggaggagaga tgagcagctg tctcctgaag aggaggagaa gcgtcgcatc cggcggggaga 720
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tcatcatctc cagaactccc agcggggctc cctgagctct caaggagatg ctgccatcac 1980
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```

<210> 68
<211> 3252
<212> DNA
<213> Homo sapiens

<220>
<221> misc_feature
<222> 779
<223> n = A,T,C or G

<400> 68
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aagtaaaaca ggatccaagc aactcgaaga acatctcaaa acatggacaa aacaatccag 420
tgggagatta tgaacatgct gatgatcaag ctgaagaaga tgctttgcaa atggcagtgg 480
gatattttga gaaaggtccc attaaagctt cacagaataa agataaaacc ttggaaaaac 540
acttgaaaac tgtggaaaat gtggcttgga agaatgggtt agcttcagaa gaaattgata 600
ttctattaaa tattgcactc agtggcaaat ttggaaatgc tgtaaacaca cggatattga 660
agtgcatgat cccagcaaca gtaaatatcag aagattctgt ggttaaggca gtctcctggc 720
tttgtgttgg caagtgttct ggtagcacca aggtactttt ttatcgttgg ctggttgcna 780
tgtttgactt cattgatcgy aaggagcaaa ttaacttgct ctatggcttc ttttttgctt 840
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tatgactcta aaagccctaa gtagttggta acttcctggg, ccttcaatca tagcaatttg 3060
atgagggaag gaaggggaga ggatttgttg ggtaatcaag acattcccgt atatgtctga 3120.
```

```
tttcatggaa ctgctctatt ttgtttgtgt gtattgtata tgtatatgtg tatgtgtgcg 3180
tgtatgtgtg tgtctgtagc ttcagttttt aagtgtaagg actaaataaa ctaactgaaa 3240
ttttactttc ag                                                      3252
```

<210> 69
<211> 756
<212> PRT
<213> Homo sapiens

<400> 69

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Met Ser Pro Gln Lys Arg Val Lys Asn Val Gln Ala Gln Asn Arg Thr
1               5                  10                 15
Ser Gln Gly Ser Ser Ser Phe Gln Thr Thr Leu Ser Ala Trp Lys Val
            20                  25                 30
Lys Gln Asp Pro Ser Asn Ser Lys Asn Ile Ser Lys His Gly Gln Asn
            35                  40                 45
Asn Pro Val Gly Asp Tyr Glu His Ala Asp Asp Gln Ala Glu Glu Asp
        50                  55                 60
Ala Leu Gln Met Ala Val Gly Tyr Phe Glu Lys Gly Pro Ile Lys Ala
65                  70                 75                 80
Ser Gln Asn Lys Asp Lys Thr Leu Glu Lys His Leu Lys Thr Val Glu
                85                  90                 95
Asn Val Ala Trp Lys Asn Gly Leu Ala Ser Glu Glu Ile Asp Ile Leu
            100                 105                110
Leu Asn Ile Ala Leu Ser Gly Lys Phe Gly Asn Ala Val Asn Thr Arg
            115                 120                125
Ile Leu Lys Cys Met Ile Pro Ala Thr Val Ile Ser Glu Asp Ser Val
        130                 135                140
Val Lys Ala Val Ser Trp Leu Cys Val Gly Lys Cys Ser Gly Ser Thr
145                 150                 155                160
Lys Val Leu Phe Tyr Arg Trp Leu Val Ala Met Phe Asp Phe Ile Asp
                165                 170                175
Arg Lys Glu Gln Ile Asn Leu Leu Tyr Gly Phe Phe Phe Ala Ser Leu
            180                 185                190
Gln Asp Asp Ala Leu Cys Pro Tyr Val Cys His Leu Leu Tyr Leu Leu
        195                 200                205
Thr Lys Lys Glu Asn Val Lys Pro Phe Arg Val Arg Lys Leu Leu Asp
    210                 215                220
Leu Gln Ala Lys Met Gly Met Gln Pro His Leu Gln Ala Leu Leu Ser
225                 230                 235                240
Leu Tyr Lys Phe Phe Ala Pro Ala Leu Ile Ser Val Ser Leu Pro Val
                245                 250                255
Arg Lys Lys Ile Tyr Leu Gln Asn Ser Glu Asn Leu Trp Lys Thr Ala
            260                 265                270
Leu Leu Ala Val Lys Gln Arg Asn Arg Gly Pro Ser Pro Glu Pro Leu
        275                 280                285
Lys Leu Met Leu Gly Pro Ala Asn Val Arg Pro Leu Lys Arg Lys Trp
    290                 295                300
Asn Ser Leu Ser Val Ile Pro Val Leu Asn Ser Ser Ser Tyr Thr Lys
305                 310                 315                320
Glu Cys Gly Lys Lys Glu Met Ser Leu Ser Asp Cys Leu Asn Arg Ser
                325                 330                335
Gly Ser Phe Pro Leu Glu Gln Leu Gln Ser Phe Pro Gln Leu Leu Gln
            340                 345                350
Asn Ile His Cys Leu Glu Leu Pro Ser Gln Met Gly Ser Val Leu Asn
        355                 360                365
Asn Ser Leu Leu Leu His Tyr Ile Asn Cys Val Arg Asp Glu Pro Val
    370                 375                380
Leu Leu Arg Phe His Tyr Trp Leu Ser Gln Thr Leu Gln Glu Glu Cys
```

```
385                    390                    395                    400
Ile Trp Tyr Lys Val Asn Asn Tyr Glu His Gly Lys Glu Phe Thr Asn
            405                    410                    415
Phe Leu Asp Thr Ile Ile Arg Ala Glu Cys Phe Leu Gln Glu Gly Tyr
            420                    425                    430
Tyr Ser Cys Glu Ala Phe Leu Tyr Lys Ser Leu Pro Leu Trp Asp Gly
            435                    440                    445
Leu Ser Cys Arg Ser Gln Phe Leu Gln Leu Val Ser Trp Ile Pro Phe
            450                    455                    460
Ser Ser Phe Ser Glu Val Lys Pro Leu Leu Phe Asp His Leu Ala Gln
465                    470                    475                    480
Leu Phe Phe Thr Ser Thr Ile Tyr Phe Lys Cys Ser Val Leu Gln Ser
                485                    490                    495
Leu Lys Glu Leu Leu Gln Asn Trp Leu Leu Trp Leu Ser Met Asp Ile
            500                    505                    510
His Met Lys Pro Val Thr Asn Ser Pro Leu Glu Thr Thr Leu Gly Gly
            515                    520                    525
Ser Met Asn Cys Val Ser Lys Leu Ile His Tyr Val Gly Trp Leu Ser
            530                    535                    540
Thr Thr Ala Met Arg Leu Glu Ser Asn Asn Thr Phe Leu Leu His Phe
545                    550                    555                    560
Ile Leu Asp Phe Tyr Glu Lys Val Cys Asp Ile Tyr Ile Asn Tyr Asp
                565                    570                    575
Leu Pro Leu Val Val Leu Phe Pro Pro Gly Ile Phe Tyr Ser Ala Leu
                580                    585                    590
Leu Ser Leu Asp Thr Ser Ile Leu Asn Gln Leu Cys Phe Ile Met His
                595                    600                    605
Arg Tyr Arg Lys Asn Leu Thr Ala Ala Lys Lys Asn Glu Leu Val Gln
            610                    615                    620
Lys Thr Lys Ser Glu Phe Asn Phe Ser Ser Lys Thr Tyr Gln Glu Phe
625                    630                    635                    640
Asn Tyr Tyr Leu Thr Ser Met Val Gly Cys Leu Trp Thr Ser Lys Pro
                645                    650                    655
Phe Ala Lys Gly Ile Tyr Ile Asp Pro Glu Ile Leu Glu Lys Thr Gly
            660                    665                    670
Val Ala Glu Tyr Lys Asn Ser Leu Asn Val Val His His Pro Ser Phe
            675                    680                    685
Leu Ser Tyr Ala Val Ser Phe Leu Leu Gln Glu Ser Pro Glu Glu Arg
            690                    695                    700
Thr Val Asn Val Ser Ser Ile Arg Gly Lys Lys Trp Ser Trp Tyr Leu
705                    710                    715                    720
Asp Tyr Leu Phe Ser Gln Gly Leu Gln Gly Leu Lys Leu Phe Ile Arg
                725                    730                    735
Ser Ser Val His His Ser Ser Ile Pro Arg Ala Glu Gly Ile Asn Cys
            740                    745                    750
Asn Asn Gln Tyr
            755
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<210> 70
<211> 1559
<212> DNA
<213> Homo sapiens

<400> 70
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cccagagtcc cttatcccta tgcccctcat ttcccctgct gtttgccect cagtctttat 120
atctcttcct tttcctcctc atcttttctc ccttcccgct tttttcctct tccttcaaag 180
tcttttcct tctctccttc ctatgctagc ctcctagctc cctcttgtgt ccctccctt   240

131

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ctgctttgtc cttttccact gtccgcactg catctgactc ctgcagagac cttgttctcc 420
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cattgggtat ggggccaggc ccctggatga atatcctgtg ggcctggttt attttctggt 1260
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```

<210> 71
<211> 338
<212> PRT
<213> Homo sapiens

<400> 71

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Met Ala Ser Ser Gly Tyr Val Leu Gln Ala Glu Leu Ser Pro Ser Thr
 1               5                   10                  15
Glu Asn Ser Ser Gln Leu Asp Phe Glu Asp Val Trp Asn Ser Ser Tyr
                20                  25                  30
Gly Val Asn Asp Ser Phe Pro Asp Gly Asp Tyr Asp Ala Asn Leu Glu
            35                  40                  45
Ala Ala Ala Pro Cys His Ser Cys Asn Leu Leu Asp Asp Ser Ala Leu
        50                  55                  60
Pro Phe Phe Ile Leu Thr Ser Val Leu Gly Ile Leu Ala Ser Ser Thr
65                  70                  75                  80
Val Leu Phe Met Leu Phe Arg Pro Leu Phe Arg Trp Gln Leu Cys Pro
                85                  90                  95
Gly Trp Pro Val Leu Ala Gln Leu Ala Val Gly Ser Ala Leu Phe Ser
            100                 105                 110
Ile Val Val Pro Val Leu Ala Pro Gly Leu Gly Ser Thr Arg Ser Ser
        115                 120                 125
Ala Leu Cys Ser Leu Gly Tyr Cys Val Trp Tyr Gly Ser Ala Phe Ala
    130                 135                 140
Gln Ala Leu Leu Leu Gly Cys His Ala Ser Leu Gly His Arg Leu Gly
145                 150                 155                 160
Ala Gly Gln Val Pro Gly Leu Thr Leu Gly Leu Thr Val Gly Ile Trp
                165                 170                 175
Gly Val Ala Ala Leu Leu Thr Leu Pro Val Thr Leu Ala Ser Gly Ala
            180                 185                 190
Ser Gly Gly Leu Cys Thr Leu Ile Tyr Ser Thr Glu Leu Lys Ala Leu
        195                 200                 205
Gln Ala Thr His Thr Val Ala Cys Leu Ala Ile Phe Val Leu Leu Pro
    210                 215                 220
Leu Gly Leu Phe Gly Ala Lys Gly Leu Lys Lys Ala Leu Gly Met Gly
225                 230                 235                 240
```

```
Pro Gly Pro Trp Met Asn Ile Leu Trp Ala Trp Phe Ile Phe Trp Trp
                245                     250             255
Pro His Gly Val Val Leu Gly Leu Asp Phe Leu Val Arg Ser Lys Leu
            260             265             270
Leu Leu Leu Ser Thr Cys Leu Ala Gln Gln Ala Leu Asp Leu Leu Leu
        275             280             285
Asn Leu Ala Glu Ala Leu Ala Ile Leu His Cys Val Ala Thr Pro Leu
    290             295             300
Leu Leu Ala Leu Phe Cys His Gln Ala Thr Arg Thr Leu Leu Pro Ser
305             310             315                     320
Leu Pro Leu Pro Glu Gly Trp Ser Ser His Leu Asp Thr Leu Gly Ser
                325                     330                 335
Lys Ser
```

<210> 72
<211> 817
<212> DNA
<213> Homo sapiens

<400> 72
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gaaccgttta ctcgctgctg tgcccatcta tcagcaggct ccgggctgaa gattgcttct 60
cttctctcct ccaaggtcta gtgacggagc ccgcgcgcgg cgccaccatg cggcagaagg 120
cggtatcgct tttcttgtgc tacctgctgc tcttcacttg cagtgggggtg gaggcaggta 180
agaaaaagtg ctcggagagc tcggacagcg gctccgggtt ctggaaggcc ctgaccttca 240
tggccgtcgg aggaggactc gcagtcgccg ggctgcccgc gctgggcttc accggcgccg 300
gcatcgcggc caactcggtg gctgcctcgc tgatgagctg gtctgcgatc ctgaatgggg 360
gcggcgtgcc cgccgggggg ctagtggcca cgctgcagag cctcggggct ggtggcagca 420
gcgtcgtcat aggtaatatt ggtgccctga tgggctacgc cacccacaag tatctcgata 480
gtgaggagga tgaggagtag ccagcagctc ccagaacctc ttcttccttc ttggcctaac 540
tcttccagtt aggatctaga actttgcctt tttttttttt tttttttttt tttgagatgg 600
gttctcacta tattgtccag gctagagtgc agtggctatt cacagatgcg aacatagtac 660
actgcagcct ccaactccta gcctcaagtg atcctcctgt ctcaacctcc caagtaggat 720
tacaagcatg cgccgacgat gcccagaatc cagaactttg tctatcactc tccccaacaa 780
cctagatgtg aaaacagaat aaacttcacc cagaaaa                            817
```

<210> 73
<211> 130
<212> PRT
<213> Homo sapiens

<400> 73
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Met Arg Gln Lys Ala Val Ser Leu Phe Leu Cys Tyr Leu Leu Leu Phe
  1               5               10              15
Thr Cys Ser Gly Val Glu Ala Gly Lys Lys Lys Cys Ser Glu Ser Ser
            20              25              30
Asp Ser Gly Ser Gly Phe Trp Lys Ala Leu Thr Phe Met Ala Val Gly
        35              40              45
Gly Gly Leu Ala Val Ala Gly Leu Pro Ala Leu Gly Phe Thr Gly Ala
    50              55              60
Gly Ile Ala Ala Asn Ser Val Ala Ala Ser Leu Met Ser Trp Ser Ala
65              70              75              80
Ile Leu Asn Gly Gly Gly Val Pro Ala Gly Gly Leu Val Ala Thr Leu
            85              90              95
Gln Ser Leu Gly Ala Gly Gly Ser Ser Val Val Ile Gly Asn Ile Gly
        100             105             110
Ala Leu Met Gly Tyr Ala Thr His Lys Tyr Leu Asp Ser Glu Glu Asp
    115             120             125
```

Glu Glu
   130

<210> 74
<211> 2861
<212> DNA
<213> Homo sapiens

<400> 74
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gccaagctgc aggggatttg ggggatgtgg gacctccaat tcccagcccc ggcttcagct 120
ctttcccagg tgttgactcc agctccagct tcagctccag ctccaggtcg ggctccagct 180
ccagccgcag cttaggcagc ggaggttctg tgtcccagtt gttttccaat ttcaccggct 240
ccgtggatga ccgtgggacc tgccagtgct ctgtttccct gccagacacc acctttcccg 300
tggacagagt ggaacgcttg gaattcacag ctcatgttct ttctcagaag tttgagaaag 360
aactttccaa agtgagggaa tatgtccaat taattagttt gtatgaaaag aaactgttaa 420
acctaactgt ccgaattgac atcatgggag aaggatacat ttcttacact gaactggact 480
tcgagctgat aaggtagaag tgaaggagat ggaaaaactg gtcatacagc tgaaggagag 540
ttttggtgga agctcagaaa ttgttgacca gctggaggtg gagataagaa atatgactct 600
cttggtagag aagcttgaga cactagacaa aaacaatgtc cttgccattc gccgagaaat 660
cgtggctctg aagaccaagc tgaaagagtg tgaggcctct aaagatcaaa acacccctgt 720
cgtccaccct cctcccactc cagggagctg tggtcatggt ggtgtggtga acatcagcaa 780
accgtctgtg gttcagctca actggagagg gtttttcttat ctatatggtg cttggggtag 840
ggattactct ccccagcatc caaacaaagg actgtattgg gtggcgccat gaatacaga 900
tgggagactg ttggagtatt atatactgta caacacactg gatgatttgc tattgtatat 960
aaatgctcga gagttgcgga tcacctatgg ccaaggtagt ggtacagcag tttacaacaa 1020
caacatgtac gtcaacatgt acacaccggg aatattgcca gagttaacct gaccaccaac 1080
acgattgctg tgactcaaac tctccctaat gctgcctata ataaccgctt ttcatatgct 1140
aatgttgctt ggcaagcata ttgactttgc tgtggatgag aatggattgt gggttatttta 1200
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gttaccttct gaattatgat ctttctgtct tgcagaagcc ccagtaagct gtttaggagt 1560
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cagatatcta agtaagtgga gaagactatt ttttcaacaa acatttttctc tttcacccta 1680
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atatattcta catctgtaaa gtgctgagtt ttatggagag aggccttttt atgcattaaa 2160
ttgtacatgg caaataaatc ccagaaggat ctgtagatga ggcacctgct ttttctttc 2220
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ctactgccta ctttaaaaaa attaatagtt ttctatggaa ctgatctaag attagaaaaa 2400
ttaatttct ttaatttcat tatgaacttt tatttacatg actctaagac tataagaaaa 2460
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tgtgcaactt atgagtgtat cagttgttgc atgtaatttt tgcctttgtt taagcctgga 2580
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tgtgcttttg aatgacttta tcatctagtc tttgtctatt tttcctttga tgttcaagtc 2760
ctagtctata ggattggcag tttaaatgct ttactccccc ttttaaaata aatgattaaa 2820
atgtgcttcg aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa a   2861

<210> 75
<211> 187
<212> PRT
<213> Homo sapiens

<400> 75

```
Met Glu Lys Leu Val Ile Gln Leu Lys Glu Ser Phe Gly Gly Ser Ser
1               5                  10                 15
Glu Ile Val Asp Gln Leu Glu Val Glu Ile Arg Asn Met Thr Leu Leu
            20                  25                 30
Val Glu Lys Leu Glu Thr Leu Asp Lys Asn Asn Val Leu Ala Ile Arg
        35                  40                 45
Arg Glu Ile Val Ala Leu Lys Thr Lys Leu Lys Glu Cys Glu Ala Ser
    50                  55                 60
Lys Asp Gln Asn Thr Pro Val Val His Pro Pro Pro Thr Pro Gly Ser
65                  70                 75                 80
Cys Gly His Gly Gly Val Val Asn Ile Ser Lys Pro Ser Val Val Gln
            85                  90                 95
Leu Asn Trp Arg Gly Phe Ser Tyr Leu Tyr Gly Ala Trp Gly Arg Asp
            100                 105                110
Tyr Ser Pro Gln His Pro Asn Lys Gly Leu Tyr Trp Val Ala Pro Leu
    115                 120                 125
Asn Thr Asp Gly Arg Leu Leu Glu Tyr Tyr Ile Leu Tyr Asn Thr Leu
    130                 135                 140
Asp Asp Leu Leu Leu Tyr Ile Asn Ala Arg Glu Leu Arg Ile Thr Tyr
145                 150                 155                160
Gly Gln Gly Ser Gly Thr Ala Val Tyr Asn Asn Asn Met Tyr Val Asn
            165                 170                 175
Met Tyr Thr Pro Gly Ile Leu Pro Glu Leu Thr
            180                 185
```

<210> 76
<211> 956
<212> DNA
<213> Homo sapiens

<400> 76

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atcaatggcc tgcgcagggt gctggatgag ctgaccctgg ccagagccga cctggagatg  120
cagattgaga acctcaagga ggagctggcc tacctgaaga agaaccacga ggaggagatg  180
aacgccctgc gaggccaggt gggtggtgag atcaatgtgg agatggacgc tgccccaggc  240
gtggacctga ccgcatcct caacgagatg cgtgaccagt atgagaagat ggcagagaag  300
aaccgcaagg atgccgagga ttggttcttc agcaagacag aggaactgaa ccgcgaggtg  360
gccaccaaca gtgagctggt gcagagtggc aagagtgaga tctcggagct ccggcgcacc  420
atgcaggcct tggagataga gctgcagtcc cagctcagca tgaaagcatc cctggagggc  480
aacctggcgg agacagagaa ccgctactgc gtgcagctgt cccagatcca ggggctgatt  540
ggcagcgtgg aggagcagct ggcccagctt cgctgcgaga tggagcagca gaaccaggaa  600
tacaaaatcc tgctggatgt gaagacgcgg ctggagcagg agattgccac ctaccgccgc  660
ctgctggagg gagaggatgc ccacctgact cagtacaaga agaaccggt gaccacccgt  720
caggtgcgta ccattgtgga gaggtccag gatggcaagg tcatctcctc ccgcgagcag  780
gtccaccaga ccacccgctg aggactcagc tacccggcc ggccacccag gaggcaggga  840
cgcagccgcc ccatctgccc cacagtctcc ggcctctcca gcctcagccc cctgcttcag  900
tcccttcccc atgcttcctt gcctgatgac aataaaagct tgttgactca gctatg      956
```

<210> 77
<211> 266
<212> PRT
<213> Homo sapiens

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<400> 77
Asp Glu Phe Arg Thr Lys Phe Glu Thr Asp Gln Ala Leu Arg Leu Ser
1               5                   10                  15
Val Glu Ala Asp Ile Asn Gly Leu Arg Arg Val Leu Asp Glu Leu Thr
            20                  25                  30
Leu Ala Arg Ala Asp Leu Glu Met Gln Ile Glu Asn Leu Lys Glu Glu
        35                  40                  45
Leu Ala Tyr Leu Lys Lys Asn His Glu Glu Glu Met Asn Ala Leu Arg
    50                  55                  60
Gly Gln Val Gly Gly Glu Ile Asn Val Glu Met Asp Ala Ala Pro Gly
65                  70                  75                  80
Val Asp Leu Ser Arg Ile Leu Asn Glu Met Arg Asp Gln Tyr Glu Lys
            85                  90                  95
Met Ala Glu Lys Asn Arg Lys Asp Ala Glu Asp Trp Phe Phe Ser Lys
            100                 105                 110
Thr Glu Glu Leu Asn Arg Glu Val Ala Thr Asn Ser Glu Leu Val Gln
        115                 120                 125
Ser Gly Lys Ser Glu Ile Ser Glu Leu Arg Arg Thr Met Gln Ala Leu
    130                 135                 140
Glu Ile Glu Leu Gln Ser Gln Leu Ser Met Lys Ala Ser Leu Glu Gly
145                 150                 155                 160
Asn Leu Ala Glu Thr Glu Asn Arg Tyr Cys Val Gln Leu Ser Gln Ile
            165                 170                 175
Gln Gly Leu Ile Gly Ser Val Glu Glu Gln Leu Ala Gln Leu Arg Cys
            180                 185                 190
Glu Met Glu Gln Gln Asn Gln Glu Tyr Lys Ile Leu Leu Asp Val Lys
        195                 200                 205
Thr Arg Leu Glu Gln Glu Ile Ala Thr Tyr Arg Arg Leu Leu Glu Gly
    210                 215                 220
Glu Asp Ala His Leu Thr Gln Tyr Lys Lys Glu Pro Val Thr Thr Arg
225             230                 235                 240
Gln Val Arg Thr Ile Val Glu Glu Val Gln Asp Gly Lys Val Ile Ser
            245                 250                 255
Ser Arg Glu Gln Val His Gln Thr Thr Arg
            260                 265


<210> 78
<211> 1689
<212> DNA
<213> Homo sapiens

<400> 78
cgggagcgtg gggtatctcg aggtgccggg ttgcaggcgc tcaggggcgc tagggtttga 60
ggcctgcttt ctgctcgcgc cagcagagca ctacctgagg cagcgaggcg cagcgagcct 120
agcctccccg cgccctgggc agtgtggcca tggagaatca ggtgttgacg ccgcatgtct 180
actgggctca gcgacaccgc gagctatatc tgcgcgtgga gctgagtgac gtacagaacc 240
ctgccatcag catcactgaa aacgtgctgc atttcaaagc tcaaggacat ggtgccaaag 300
gagacaatgt ctatgaattt cacctggagt tcttagacct tgtgaaacca gagcctgttt 360
acaaactgac ccagaggcag gtaaacatta cagtacagaa gaaagtgagt cagtggtggg 420
agagactcac aaagcaggaa aagcgaccac tgtttttggc tcctgacttt gatcgttggc 480
tggatgaatc tgatgcggaa atggagctca gagctaagga agaagagcgc ctaaataaac 540
tccgactgga aagcgaaggc tctcctgaaa ctcttacaaa cttaggaaaa ggatacctgt 600
ttatgtataa tcttgtgcaa ttcttgggat tctcctggat ctttgtcaac ctgactgtgc 660
gattctgtat cttgggaaaa gagtcctttt atgacacatt ccatactgtg gctgacatga 720
tgtatttctg ccagatgctg gcagttgtgg aaactatcaa tgcagcaatt ggagtcacta 780
cgtcaccggt gctgccttct ctgatccagc ttcttggaag aaattttatt ttgtttatca 840
tctttggcac catggaagaa atgcagaaca aagctgtggt tttctttgtg ttttatttgt 900
```

```
ggagtgcaat tgaaattttc aggtactctt tctacatgct gacgtgcatt gacatggatt 960
ggaaggtgct cacatggctt cgttacactc tgtggattcc cttatatcca ctgggatgtt 1020
tggtggaagc tgtctcagtg attcagtcca ttccaatatt caatgagacc ggacgattca 1080
gtttcacatt gccatatcca gtgaaaatca aagttagatt ttcctttttt cttcagattt 1140
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gacggcgcta tggaaaaaaa agaaaaagat ccactaaaaa gaaagattta gatggcttct 1260
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cctgaattta ctgttatctt attgtagtac ttgcatgaca tggattcctg atatctgatg 1440
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cctacaatgt actgacagtt cttacagttg aggatttggt tcttttcagc taattgcttg 1620
gtggattaaa aaaagcaaga ctaatgtcaa ctctaatgga aggctggtta aaagtggact 1680
caggcaagg                                                        1689
```

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<210> 79
<211> 373
<212> PRT
<213> Homo sapiens
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<400> 79
Met Glu Asn Gln Val Leu Thr Pro His Val Tyr Trp Ala Gln Arg His
1               5                   10                  15
Arg Glu Leu Tyr Leu Arg Val Glu Leu Ser Asp Val Gln Asn Pro Ala
            20                  25                  30
Ile Ser Ile Thr Glu Asn Val Leu His Phe Lys Ala Gln Gly His Gly
        35                  40                  45
Ala Lys Gly Asp Asn Val Tyr Glu Phe His Leu Glu Phe Leu Asp Leu
    50                  55                  60
Val Lys Pro Glu Pro Val Tyr Lys Leu Thr Gln Arg Gln Val Asn Ile
65                  70                  75                  80
Thr Val Gln Lys Lys Val Ser Gln Trp Trp Glu Arg Leu Thr Lys Gln
                85                  90                  95
Glu Lys Arg Pro Leu Phe Leu Ala Pro Asp Phe Asp Arg Trp Leu Asp
            100                 105                 110
Glu Ser Asp Ala Glu Met Glu Leu Arg Ala Lys Glu Glu Glu Arg Leu
        115                 120                 125
Asn Lys Leu Arg Leu Glu Ser Glu Gly Ser Pro Glu Thr Leu Thr Asn
    130                 135                 140
Leu Arg Lys Gly Tyr Leu Phe Met Tyr Asn Leu Val Gln Phe Leu Gly
145                 150                 155                 160
Phe Ser Trp Ile Phe Val Asn Leu Thr Val Arg Phe Cys Ile Leu Gly
                165                 170                 175
Lys Glu Ser Phe Tyr Asp Thr Phe His Thr Val Ala Asp Met Met Tyr
            180                 185                 190
Phe Cys Gln Met Leu Ala Val Val Glu Thr Ile Asn Ala Ala Ile Gly
        195                 200                 205
Val Thr Thr Ser Pro Val Leu Pro Ser Leu Ile Gln Leu Leu Gly Arg
    210                 215                 220
Asn Phe Ile Leu Phe Ile Ile Phe Gly Thr Met Glu Glu Met Gln Asn
225                 230                 235                 240
Lys Ala Val Val Phe Phe Val Phe Tyr Leu Trp Ser Ala Ile Glu Ile
                245                 250                 255
Phe Arg Tyr Ser Phe Tyr Met Leu Thr Cys Ile Asp Met Asp Trp Lys
            260                 265                 270
Val Leu Thr Trp Leu Arg Tyr Thr Leu Trp Ile Pro Leu Tyr Pro Leu
        275                 280                 285
Gly Cys Leu Val Glu Ala Val Ser Val Ile Gln Ser Ile Pro Ile Phe
    290                 295                 300
```

Asn Glu Thr Gly Arg Phe Ser Phe Thr Leu Pro Tyr Pro Val Lys Ile
305   310   315   320
Lys Val Arg Phe Ser Phe Phe Leu Gln Ile Tyr Leu Ile Met Ile Phe
                325   330   335
Leu Gly Leu Tyr Ile Asn Phe Arg His Leu Tyr Lys Gln Arg Arg Arg
            340   345   350
Arg Tyr Gly Lys Lys Arg Lys Arg Ser Thr Lys Lys Lys Asp Leu Asp
        355   360   365
Gly Phe Leu Pro Val
        370

<210> 80
<211> 1824
<212> DNA
<213> Homo sapiens

<400> 80
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ggacagtgtc ccgctgcggc tccgcggcga tggccaccaa gatcgacaaa gaggcttgcc 180
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cagcccccac cttcccgctt cccttagaac cctgccgtgt cctatctcat gacgctcatg 780
gaacctcttt ctttgatctt cttttctttt tctcccctc ttttttgttc taaagaaaag 840
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tcatgagcac atttgatata gctctttttc tgtttttcct tgctcatttc gttttgggga 1740
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ataaatgttt tttttgtttt tcag   1824

<210> 81
<211> 142
<212> PRT
<213> Homo sapiens

<400> 81
Met Ala Thr Lys Ile Asp Lys Glu Ala Cys Arg Ala Ala Tyr Asn Leu
  1   5   10   15
Val Arg Asp Asp Gly Ser Ala Val Ile Trp Val Thr Phe Lys Tyr Asp

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             35                  40                  45
Gly Ile Ser Arg Arg Met Pro Pro Ala Asn Leu Pro Ser Leu Lys
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Ala Glu Asn Lys Gly Asn Asp Pro Asn Val Asn Ile Val Pro Lys Asp
65                  70                  75                  80
Gly Thr Gly Trp Ala Ser Lys Gln Glu Gln His Glu Glu Glu Lys Thr
                 85                  90                  95
Pro Glu Val Pro Pro Ala Gln Pro Lys Pro Gly Val Ala Ala Pro Pro
                100                 105                 110
Glu Val Ala Pro Ala Pro Lys Ser Trp Ala Ser Asn Lys Gln Gly Gly
            115                 120                 125
Gln Gly Asp Gly Ile Gln Val Asn Ser Gln Phe Gln Gln Glu Phe Pro
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Ser Leu Gln Ala Ala Gly Asp Gln Glu Lys Lys Glu Lys Glu Thr Asn
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Asp Asp Asn Tyr Gly Pro Gly Pro Ser Leu Arg Pro Pro Asn Val Ala
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Cys Trp Arg Asp Gly Gly Lys Ala Ala Gly Ser Pro Ser Ser Ser Asp
            180                 185                 190
Gln Asp Glu Lys Leu Pro Gly Gln Asp Glu Ser Thr Ala Gly Thr Ser
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Pro Pro Gln Ala Lys Leu Asn Gly Gln Gln Ala Ala Leu Ala Ser Gln
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Gln Gly Ser Lys Ala Ser Glu Asn Asn Glu Asn Lys Lys Glu Thr Asp
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Glu Val Ser Asn Thr Lys Ser Ser Ser Gln Ile Pro Ala Gln Pro Ser
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Val Ala Lys Val Pro Tyr Gly Lys Gly Pro Ser Phe Asn Gln Glu Arg
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Gly Thr Ser Ser His Leu Pro Pro Pro Lys Leu Leu Ala Gln Gln
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                980                   985                   990
Trp Gly Pro Arg Pro Ser Ser Asn Arg Arg Glu Glu Val Asn Asp Arg
    995                   1000                  1005
Pro Val Arg Arg Ser Gly Pro Ile Lys Lys Pro Val Leu Arg Asp Met
    1010                  1015                  1020
Lys Glu Glu Arg Glu Gln Arg Lys Glu Lys Glu Gly Glu Lys Ala Glu
1025                  1030                  1035                  1040
Lys Val Thr Glu Lys Val Val Val Lys Pro Glu Lys Thr Glu Lys Lys
                1045                  1050                  1055
Asp Leu Pro Pro Pro Pro Pro Pro Gln Pro Pro Ala Pro Ile Gln
                1060                  1065                  1070
Pro Gln Ser Val Pro Pro Pro Ile Gln Pro Glu Ala Glu Lys Phe Pro
    1075                  1080                  1085
Ser Thr Glu Thr Ala Thr Leu Ala Gln Lys Pro Ser Gln Asp Thr Glu
    1090                  1095                  1100
Lys Pro Leu Glu Pro Val Ser Thr Val Gln Val Glu Pro Ala Val Lys
1105                  1110                  1115                  1120
Thr Val Asn Gln Gln Thr Met Ala Ala Pro Val Val Lys Glu Glu Lys
                1125                  1130                  1135
Gln Pro Glu Lys Val Ile Ser Lys Asp Leu Val Ile Glu Arg Pro Arg
                1140                  1145                  1150
Pro Asp Ser Arg Pro Ala Val Lys Lys Glu Ser Thr Leu Pro Pro Arg
                1155                  1160                  1165
Thr Tyr Trp Lys Glu Ala Arg Glu Arg Asp Trp Phe Pro Asp Gln Gly
    1170                  1175                  1180
Tyr Arg Gly Arg Gly Arg Gly Glu Tyr Tyr Ser Arg Gly Arg Ser Tyr
1185                  1190                  1195                  1200
Arg Gly Ser Tyr Gly Gly Arg Gly Arg Gly Arg Gly His Thr Arg
                1205                  1210                  1215
Asp Tyr Pro Gln Tyr Arg Asp Asn Lys Pro Arg Ala Glu His Ile Pro
                1220                  1225                  1230
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144·

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Ser Gly Pro Leu Arg Gln Arg Glu Glu Ser Glu Thr Arg Ser Glu Ser
        1235            1240                1245
Ser Asp Phe Glu Val Val Pro Lys Arg Arg Arg Gln Arg Gly Ser Glu
        1250            1255                1260
Thr Asp Thr Asp Ser Glu Ile His Glu Ser Ala Ser Asp Lys Asp Ser
1265            1270                1275                1280
Leu Ser Lys Gly Lys Leu Pro Lys Arg Glu Glu Arg Pro Glu Asn Lys
            1285            1290                1295
Lys Pro Val Lys Pro His Ser Ser Phe Lys Pro Asp Asn His Val Arg
            1300            1305                1310
Ile Asp Asn Arg Leu Leu Glu Lys Pro Tyr Val Arg Asp Asp Asp Lys
            1315            1320                1325
Ala Lys Pro Gly Phe Leu Pro Lys Gly Glu Pro Thr Arg Arg Gly Arg
        1330            1335                1340
Gly Gly Thr Phe Arg Arg Gly Gly Arg Asp Pro Gly Gly Arg Pro Ser
1345            1350                1355                1360
Arg Pro Ser Thr Leu Arg Arg Pro Ala Tyr Arg Asp Asn Gln Trp Asn
            1365            1370                1375
Pro Arg Gln Ser Glu Val Pro Lys Pro Glu Asp Gly Glu Pro Pro Arg
            1380            1385                1390
Arg His Glu Gln Phe Ile Pro Ile Ala Ala Asp Lys Arg Pro Pro Lys
            1395            1400                1405
Phe Glu Arg Lys Phe Asp Pro Ala Arg Glu Arg Pro Arg Arg Gln Arg
            1410            1415                1420
Pro Thr Arg Pro Pro Arg Gln Asp Lys Pro Pro Arg Phe Arg Arg Leu
1425            1430                1435                1440
Arg Glu Arg Glu Ala Ala Ser Lys Ser Asn Glu Val Val Ala Val Pro
            1445            1450                1455
Thr Asn Gly Thr Val Asn Asn Val Ala Gln Glu Pro Val Asn Thr Leu
            1460            1465                1470
Gly Asp Ile Ser Gly Asn Lys Thr Pro Asp Leu Ser Asn Gln Asn Ser
            1475            1480                1485
Ser Asp Gln Ala Asn Glu Glu Trp Glu Thr Ala Ser Glu Ser Ser Asp
        1490            1495                1500
Phe Asn Glu Arg Arg Glu Arg Asp Glu Lys Lys Asn Ala Asp Leu Asn
1505            1510                1515                1520
Ala Gln Thr Val Val Lys Val Gly Glu Asn Val Leu Pro Pro Lys Arg
            1525            1530                1535
Glu Ile Ala Lys Arg Ser Phe Ser Ser Gln Arg Pro Val Asp Arg Gln
            1540            1545                1550
Asn Arg Arg Gly Asn Asn Gly Pro Pro Lys Ser Gly Arg Asn Phe Ser
            1555            1560                1565
Gly Pro Arg Asn Glu Arg Arg Ser Gly Pro Pro Ser Lys Ser Gly Lys
            1570            1575                1580
Arg Gly Pro Phe Asp Asp Gln Pro Ala Gly Thr Thr Gly Val Asp Leu
1585            1590                1595                1600
Ile Asn Gly Ser Ser Ala His His Gln Glu Gly Val Pro Asn Gly Thr
            1605            1610                1615
Gly Gln Lys Asn Ser Lys Asp Ser Thr Gly Lys Lys Arg Glu Asp Pro
        1620            1625                1630
Lys Pro Gly Pro Lys Lys Pro Lys Glu Lys Val Asp Ala Leu Ser Gln
            1635            1640                1645
Phe Asp Leu Asn Asn Tyr Ala Ser Val Val Ile Ile Asp Asp His Pro
        1650            1655                1660
Glu Val Thr Val Ile Glu Asp Pro Gln Ser Asn Leu Asn Asp Asp Gly
1665            1670                1675                1680
Phe Thr Glu Val Val Ser Lys Lys Gln Gln Lys Arg Leu Gln Asp Glu
            1685            1690                1695
Glu Arg Arg Lys Lys Glu Glu Gln Val Ile Gln Val Trp Asn Lys Lys
```

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               1700                1705                1710
Asn Ala Asn Glu Lys Gly Arg Ser Gln Thr Ser Lys Leu Pro Pro Arg
        1715                1720                1725
Phe Ala Lys Lys Gln Ala Thr Gly Ile Gln Gln Ala Gln Ser Ser Ala
    1730                1735                1740
Ser Val Pro Pro Leu Ala Ser Ala Pro Leu Pro Pro Ser Thr Ser Ala
1745                1750                1755                1760
Ser Val Pro Ala Ser Thr Ser Ala Pro Leu Pro Ala Thr Leu Thr Pro
            1765                1770                1775
Val Pro Ala Ser Thr Ser Ala Pro Val Pro Ala Ser Thr Leu Ala Pro
        1780                1785                1790
Val Leu Ala Ser Thr Ser Ala Pro Val Pro Ala Ser Pro Leu Ala Pro
        1795                1800                1805
Val Ser Ala Ser Ala Ser Val Ser Ala Ser Val Pro Ala Ser Thr Ser
    1810                1815                1820
Ala Ala Ala Ile Thr Ser Ser Ser Ala Pro Ala Ser Ala Pro Ala Pro
1825                1830                1835                1840
Thr Pro Ile Leu Ala Ser Val Ser Thr Pro Ala Ser Val Thr Ile Leu
            1845                1850                1855
Ala Ser Ala Ser Ile Pro Ile Leu Ala Ser Ala Leu Ala Ser Thr Ser
        1860                1865                1870
Ala Pro Thr Pro Ala Pro Ala Ala Ser Ser Pro Ala Ala Pro Val Ile
        1875                1880                1885
Thr Ala Pro Thr Ile Pro Ala Ser Ala Pro Thr Ala Ser Val Pro Leu
    1890                1895                1900
Ala Pro Ala Ser Ala Ser Ala Pro Ala Pro Ala Pro Thr Pro Val Ser
1905                1910                1915                1920
Ala Pro Asn Pro Ala Pro Pro Ala Pro Ala Gln Thr Gln Ala Gln Thr
            1925                1930                1935
His Lys Pro Val Gln Asn Pro Leu Gln Thr Thr Ser Gln Ser Ser Lys
        1940                1945                1950
Gln Pro Pro Pro Ser Ile Arg Leu Pro Ser Ala Gln Thr Pro Asn Gly
    1955                1960                1965
Thr Asp Tyr Val Ala Ser Gly Lys Ser Ile Gln Thr Pro Gln Ser His
        1970                1975                1980
Gly Thr Leu Thr Ala Glu Leu Trp Asp Asn Lys Val Ala Pro Pro Ala
1985                1990                1995                2000
Val Leu Asn Asp Ile Ser Lys Lys Leu Gly Pro Ile Ser Pro Pro Gln
            2005                2010                2015
Pro Pro Ser Val Ser Ala Trp Asn Lys Pro Leu Thr Ser Phe Gly Ser
        2020                2025                2030
Ala Pro Ser Ser Glu Gly Ala Lys Asn Gly Gln Glu Ser Gly Leu Glu
        2035                2040                2045
Ile Gly Thr Asp Thr Ile Gln Phe Gly Ala Pro Ala Ser Asn Gly Asn
    2050                2055                2060
Glu Asn Glu Val Val Pro Val Leu Ser Glu Lys Ser Ala Asp Lys Ile
2065                2070                2075                2080
Pro Glu Pro Lys Glu Gln Arg Gln Lys Gln Pro Arg Ala Gly Pro Ile
            2085                2090                2095
Lys Ala Gln Lys Leu Pro Asp Leu Ser Pro Val Glu Asn Lys Glu His
        2100                2105                2110
Lys Pro Gly Pro Ile Gly Lys Glu Arg Ser Leu Lys Asn Arg Lys Val
        2115                2120                2125
Lys Asp Ala Gln Gln Val Glu Pro Glu Gly Gln Glu Lys Pro Ser Pro
    2130                2135                2140
Ala Thr Val Arg Ser Thr Asp Pro Val Thr Thr Lys Glu Thr Lys Ala
2145                2150                2155                2160
Val Ser Glu Met Ser Thr Glu Ile Gly Thr Met Ile Ser Val Ser Ser
            2165                2170                2175
```

Ala Glu Tyr Gly Thr Asn Ala Lys Glu Ser Val Thr Asp Tyr Thr Thr
            2180   2185   2190
Pro Ser Ser Ser Leu Pro Asn Thr Val Ala Thr Asn Asn Thr Lys Met
            2195   2200   2205
Glu Asp Thr Leu Val Asn Asn Val Pro Leu Pro Asn Thr Leu Pro Leu
    2210   2215   2220
Pro Lys Arg Glu Thr Ile Gln Gln Ser Ser Ser Leu Thr Ser Val Pro
2225   2230   2235   2240
Pro Thr Thr Phe Ser Leu Thr Phe Lys Met Glu Ser Ala Arg Lys Ala
            2245   2250   2255
Trp Glu Asn Ser Pro Asn Val Arg Glu Lys Gly Ser Pro Val Thr Ser
            2260   2265   2270
Thr Ala Pro Pro Ile Ala Thr Gly Val Ser Ser Ser Ala Ser Gly Pro
            2275   2280   2285
Ser Thr Ala Asn Tyr Asn Ser Phe Ser Ser Ala Ser Met Pro Gln Ile
            2290   2295   2300
Pro Val Ala Ser Val Thr Pro Thr Ala Ser Leu Ser Gly Ala Gly Thr
2305   2310   2315   2320
Tyr Thr Thr Ser Ser Leu Ser Thr Lys Ser Thr Thr Thr Ser Asp Pro
            2325   2330   2335
Pro Asn Ile Cys Lys Val Lys Pro Gln Gln Leu Gln Thr Ser Ser Leu
            2340   2345   2350
Pro Ser Ala Ser His Phe Ser Gln Leu Ser Cys Met Pro Ser Leu Ile
            2355   2360   2365
Ala Gln Gln Gln Gln Asn Pro Gln Val Tyr Val Ser Gln Ser Ala Ala
            2370   2375   2380
Ala Gln Ile Pro Ala Phe Tyr Met Asp Thr Ser His Leu Phe Asn Thr
2385   2390   2395   2400
Gln His Ala Arg Leu Ala Pro Pro Ser Leu Ala Gln Gln Gln Gly Phe
            2405   2410   2415
Gln Pro Gly Leu Ser Gln Pro Thr Ser Val Gln Gln Ile Pro Ile Pro
            2420   2425   2430
Ile Tyr Ala Pro Leu Gln Gly Gln His Gln Ala Gln Leu Ser Leu Gly
            2435   2440   2445
Ala Gly Pro Ala Val Ser Gln Ala Gln Glu Leu Phe Ser Ser Ser Leu
    2450   2455   2460
Gln Pro Tyr Arg Ser Gln Pro Ala Phe Met Gln Ser Ser Leu Ser Gln
2465   2470   2475   2480
Pro Ser Val Val Leu Ser Gly Thr Ala Ile His Asn Phe Pro Thr Val
            2485   2490   2495
Gln His Gln Glu Leu Ala Lys Ala Gln Ser Gly Leu Ala Phe Gln Gln
            2500   2505   2510
Thr Ser Asn Thr Gln Pro Ile Pro Ile Leu Tyr Glu His Gln Leu Gly
            2515   2520   2525
Gln Ala Ser Gly Leu Gly Gly Ser Gln Leu Ile Asp Thr His Leu Leu
    2530   2535   2540
Gln Ala Arg Ala Asn Leu Thr Gln Ala Ser Asn Leu Tyr Ser Gly Gln
2545   2550   2555   2560
Val Gln Gln Pro Gly Gln Thr Asn Phe Tyr Asn Thr Ala Gln Ser Pro
            2565   2570   2575
Ser Ala Leu Gln Gln Val Thr Val Pro Leu Pro Ala Ser Gln Leu Ser
            2580   2585   2590
Leu Pro Asn Phe Gly Ser Thr Gly Gln Pro Leu Ile Ala Leu Pro Gln
            2595   2600   2605
Thr Leu Gln Pro Pro Leu Gln His Thr Thr Pro Gln Ala Gln Ala Gln
    2610   2615   2620
Ser Leu Ser Arg Pro Ala Gln Val Ser Gln Pro Phe Arg Gly Leu Ile
2625   2630   2635   2640
Pro Ala Gly Thr Gln His Ser Met Ile Ala Thr Thr Gly Lys Met Ser

```
                     2645                 2650                 2655
Glu Met Glu Leu Lys Ala Phe Gly Ser Gly Ile Asp Ile Lys Pro Gly
             2660                 2665                 2670
Thr Pro Pro Ile Ala Gly Arg Ser Thr Thr Pro Thr Ser Ser Pro Ser
             2675                 2680                 2685
Gly Leu Leu Leu Gln Val Arg Thr Ala Ser Pro Ala Lys
        2690                 2695                 2700
```

<210> 84
<211> 597
<212> DNA
<213> Homo sapiens

<400> 84

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agctgaagtt gaggatctct tactctctaa gccacggaat taacccgagc aggcatggag 60
gcctctgctc tcacctcatc agcagtgacc agtgtggcca aagtggtcag ggtggcctct 120
ggctctgccg tagtttgcc cctggccagg attgctacag ttgtgattgg aggagttgtg 180
gccatggcgg ctgtgcccat ggtgctcagt gccatgggct tcactgcggc gggaatcgcc 240
tcgtcctcca tagcagccaa gatgatgtcc gcggcggcca ttgccaatgg gggtggagtt 300
gcctcgggca gccttgtggg tactctgcag tcactgggag caactggact ctccggattg 360
accaagttca tcctgggctc cattgggtct gccattgcgg ctgtcattgc gaggttctac 420
tagctccctg cccctcgccc tgcagagaag agaaccatgc caggggagaa ggcacccagc 480
catcctgacc cagcgaggag ccaactatcc caaatatacc tgggtgaaat ataccaaatt 540
ctgcatctcc agaggaaat aagaaataaa gatgaattgt tgcaactctt aaaaaaa    597
```

<210> 85
<211> 122
<212> PRT
<213> Homo sapiens

<400> 85

```
Met Glu Ala Ser Ala Leu Thr Ser Ser Ala Val Thr Ser Val Ala Lys
  1               5                 10                  15
Val Val Arg Val Ala Ser Gly Ser Ala Val Val Leu Pro Leu Ala Arg
            20                  25                  30
Ile Ala Thr Val Val Ile Gly Gly Val Val Ala Met Ala Ala Val Pro
            35                  40                  45
Met Val Leu Ser Ala Met Gly Phe Thr Ala Ala Gly Ile Ala Ser Ser
      50                  55                  60
Ser Ile Ala Ala Lys Met Met Ser Ala Ala Ala Ile Ala Asn Gly Gly
65                  70                  75                  80
Gly Val Ala Ser Gly Ser Leu Val Gly Thr Leu Gln Ser Leu Gly Ala
                85                  90                  95
Thr Gly Leu Ser Gly Leu Thr Lys Phe Ile Leu Gly Ser Ile Gly Ser
            100                 105                 110
Ala Ile Ala Ala Val Ile Ala Arg Phe Tyr
            115                 120
```

<210> 86
<211> 1032
<212> DNA
<213> Homo sapiens

<400> 86

```
ggagggtggg cagcactcgc tttattgtcc agcattccac atggatagtc gccacacctt 60
tgcccctgct gcgatgaccc tgtcgccact tctgctgttc ctgccaccgc tgctgctgct 120
gctggacgtc cccacggcgg cggtgcaggc gtccctctg caagcgttag acttctttgg 180
```

```
gaatgggcca ccagttaact acaagacagg caatctatac ctgcgggggc ccctgaagaa 240
gtccaatgca ccgcttgtca atgtgaccct ctactatgaa gcactgtgcg gtggctgccg 300
agccttcctg atccgggagc tcttcccaac atggctgttg gtcatggaga tcctcaatgt 360
cacgtcggtg ccctacggaa acgcacagga acaaaatgtc agtggcaggt gggagttcaa 420
gtgccagctt ggagaagagg agtgcaaatt caacaaggtg gaggcctgcg tgttggatga 480
acttgacatg gagctagcct tcctgaccat gtctggcatg gcatggaaga gtttgaggac 540
atggagagaa gtctgccact atgcctgcag ctctacgccc cagggctgtc gccagaacta 600
tcatggagtg tgcaatgggg gaccgcggca tgcagctcat gcacgccaac gcccagcgga 660
cagatgctct ccagccaccg cacgagtatg tgccctgggt caccgtcaat gggaaaccct 720
tggaagatca gacccagctc cttacccttg tctgccagtt gtaccagggc aagaagccgg 780
atgtctgccc ttcctcaacc agctccctcc ggagtgtttg cttcgagtgt tggccggtgg 840
gctgcggaga gctcatggaa ggcgagtggg aactcggctg cctgcctttt tttctgatcc 900
agaccctcgg cacctgctac ttaccaactg gaaaatttta tgcatcccat gaagcccaga 960
tacacaaaat tccacccta gatcaagaat cctgctccac taagaatggt gctaaagtaa 1020
aactagttta at                                                   1032
```

```
<210> 87
<211> 303
<212> PRT
<213> Homo sapiens
```

```
<400> 87
Met Asp Ser Arg His Thr Phe Ala Pro Ala Ala Met Thr Leu Ser Pro
 1               5                  10                  15
Leu Leu Leu Phe Leu Pro Pro Leu Leu Leu Leu Asp Val Pro Thr
            20                  25                  30
Ala Ala Val Gln Ala Ser Pro Leu Gln Ala Leu Asp Phe Phe Gly Asn
            35                  40                  45
Gly Pro Pro Val Asn Tyr Lys Thr Gly Asn Leu Tyr Leu Arg Gly Pro
    50                  55                  60
Leu Lys Lys Ser Asn Ala Pro Leu Val Asn Val Thr Leu Tyr Tyr Glu
65                  70                  75                  80
Ala Leu Cys Gly Gly Cys Arg Ala Phe Leu Ile Arg Glu Leu Phe Pro
                85                  90                  95
Thr Trp Leu Leu Val Met Glu Ile Leu Asn Val Thr Ser Val Pro Tyr
            100                 105                 110
Gly Asn Ala Gln Glu Gln Asn Val Ser Gly Arg Trp Glu Phe Lys Cys
            115                 120                 125
Gln Leu Gly Glu Glu Glu Cys Lys Phe Asn Lys Val Glu Ala Cys Val
        130                 135                 140
Leu Asp Glu Leu Asp Met Glu Leu Ala Phe Leu Thr Met Ser Gly Met
145                 150                 155                 160
Ala Trp Lys Ser Leu Arg Thr Trp Arg Glu Val Cys His Tyr Ala Cys
                165                 170                 175
Ser Ser Thr Pro Gln Gly Cys Arg Gln Asn Tyr His Gly Val Cys Asn
            180                 185                 190
Gly Gly Pro Arg His Ala Ala His Ala Arg Gln Arg Pro Ala Asp Arg
            195                 200                 205
Cys Ser Pro Ala Thr Ala Arg Val Cys Ala Leu Gly His Arg Gln Trp
        210                 215                 220
Glu Thr Leu Gly Arg Ser Asp Pro Ala Pro Tyr Pro Cys Leu Pro Val
225                 230                 235                 240
Val Pro Gly Gln Glu Ala Gly Cys Leu Pro Phe Leu Asn Gln Leu Pro
                245                 250                 255
Pro Glu Cys Leu Leu Arg Val Leu Ala Gly Gly Leu Arg Arg Ala His
            260                 265                 270
Gly Arg Arg Val Gly Thr Arg Leu Pro Ala Phe Phe Ser Asp Pro Asp
            275                 280                 285
Pro Arg His Leu Leu Leu Thr Asn Trp Lys Ile Leu Cys Ile Pro
```

290   295   300

<210> 88
<211> 905
<212> DNA
<213> Homo sapiens

<400> 88
caacacaggg gcagtctcca ggacctccac accattaaca agatgagcct tgtgctccct 60
tgggctctag agaggaagcc cctctgagcc ctcagcccct ctttcctccc tctcctaaag 120
taatttgatc ctcaggaatt tgttctgccc tcatctggcc ctggccagct ctgcatttga 180
caaatgccag gaagaggaaa ctgttgagaa aacggaacta ctggggaaag ggagggctca 240
ctgagaacca tcccggtaac ccgaccgccg ctggtcacca tgaaccacat tgtgcaaacc 300
ttctctcctg tcaacagcgg ccagcctccc aactacgaga tgctcaagga ggagcaggaa 360
gtggctatgc tgggggggcc ccacaaccct gctcccccga cgtccaccgt gatccacatc 420
cgcagcgaga cctccgtgcc tgaccatgtc gtctggtccc tgttcaacac cctcttcatg 480
aacacctgct gcctgggctt catagcattc gcctactccg tgaagtctag ggacaggaag 540
atggttggcg acgtgaccgg ggcccaggcc tatgcctcca ccgccaagtg cctgaacatc 600
tgggccctga ttttgggcat cttcatgacc attctgctcg tcatcatccc agtgttggtc 660
gtccaggccc agcgatagat caggaggcat cattgaggcc aggagctctg cccgtgacct 720
gtatcccacg tactctatct tccattcctc gccctgcccc cagaggccag gagctctgcc 780
cttgacctgt attccactta ctccaccttc cattcctcgc cctgtcccca cagccgagtc 840
ctgcatcagc cctttatcct cacacgcttt tctacaatgg cattcaataa agtgtatatg 900
tttct   905

<210> 89
<211> 132
<212> PRT
<213> Homo sapiens

<400> 89
Met Asn His Ile Val Gln Thr Phe Ser Pro Val Asn Ser Gly Gln Pro
1   5   10   15
Pro Asn Tyr Glu Met Leu Lys Glu Glu Gln Glu Val Ala Met Leu Gly
            20   25   30
Gly Pro His Asn Pro Ala Pro Pro Thr Ser Thr Val Ile His Ile Arg
        35   40   45
Ser Glu Thr Ser Val Pro Asp His Val Val Trp Ser Leu Phe Asn Thr
    50   55   60
Leu Phe Met Asn Thr Cys Cys Leu Gly Phe Ile Ala Phe Ala Tyr Ser
65   70   75   80
Val Lys Ser Arg Asp Arg Lys Met Val Gly Asp Val Thr Gly Ala Gln
                85   90   95
Ala Tyr Ala Ser Thr Ala Lys Cys Leu Asn Ile Trp Ala Leu Ile Leu
            100   105   110
Gly Ile Phe Met Thr Ile Leu Leu Val Ile Ile Pro Val Leu Val Val
            115   120   125
Gln Ala Gln Arg
        130

<210> 90
<211> 2499
<212> DNA
<213> Homo sapiens

<400> 90
agatgcgagc actgcggctg ggcgctgagg atcagccgct tcctgcctgg attccacagc 60

150

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ttcgcgccgt gtactgtcgc cccatccctg cgcgcccagc ctgccaagca gcgtgccccg 120
gttgcaggcg tcatgcagcg ggcgcgaccc acgctctggg ccgctgcgct gactctgctg 180
gtgctgctcc gcgggccgcc ggtggcgcgg gctggcgcga gctcgggggg cttgggtccc 240
gtggtgcgct gcgagccgtg cgacgcgcgt gcactggccc agtgcgcgcc tccgcccgcc 300
gtgtgcgcgg agctggtgcg cgagccgggc tgcggctgct gcctgacgtg cgcactgagc 360
gagggccagc cgtgcggcat ctacaccgag cgctgtggct ccggccttcg ctgccagccg 420
tcgcccgacg aggcgcgacc gctgcaggcg ctgctggacg gccgcgggct ctgcgtcaac 480
gctagtgccg tcagccgcct gcgcgcctac ctgctgccag cgccgccagc tccaggaaat 540
gctagtgagt cggaggaaga ccgcagcgcc ggcagtgtgg agagcccgtc cgtctccagc 600
acgcaccggg tgtctgatcc caagttccac cccctccatt caaagataat catcatcaag 660
aaagggcatg ctaaagacag ccagcgctac aaagttgact acgagtctca gagcacagat 720
acccagaact tctcctccga gtccaagcgg gagacagaat atggtccctg ccgtagagaa 780
atggaagaca cactgaatca cctgaagttc ctcaatgtgc tgagtcccag gggtgtacac 840
attcccaact gtgacaagaa gggatttttat aagaaaaagc agtgtcgccc ttccaaaggc 900
aggaagcggg gcttctgctg gtgtgtggat aagtatgggc agcctctccc aggctacacc 960
accaagggga aggaggacgt gcactgctac agcatgcaga gcaagtagac gcctgccgca 1020
aggttaatgt ggagctcaaa tatgccttat tttctacaaa agactgccaa ggacatgacc 1080
agcagctggc tacagcctcg atttatattt ctgtttgtgg tgaactgatt ttttttaaac 1140
caaagtttag aaagaggttt ttgaaatgcc tatggtttct ttgaatggta aacttgagca 1200
tcttttcact ttccagtagt cagcaaagag cagtttgaat tttcttgtcg cttcctatca 1260
aaatatctag agactcgagc acagcaccca gacttcatgc gcccgtggaa tgctcaccac 1320
atgttggtcg aagcggccga ccactgactt tgtgacttag gcggctgtgt tgcctatgta 1380
gagaacacgc ttcaccccca ctccctgtac agtgcgcaca ggctttatcg agaataggaa 1440
aacctttaaa ccccggtcat ccggacatcc caacgcatgc tcctggagct cacagccttc 1500
tgtggtgtca tttctgaaac aagggcgtgg atccctcaac ccagaagagt gtttatgtct 1560
tcaagtgacc tgtactgctt ggggactatt tgagaaaata aggtggagtc ctacttgttt 1620
cacaaatatg tatctaagaa tgttctaggg cactctggga acctataaag gcaggtattt 1680
cgggccctcc tcttcaggaa tcttcctgaa gacatggccc agtcgaaggc ccaggatggc 1740
ttttgctgcg gccccgtggg gtaggaggga cagagagaca gggagagtca gcctccacat 1800
tcagaggcat cacaagtaat ggcacaattc ttcggatgac tgcagaaaat agtgttttgt 1860
agttcaacaa ctcaagacga agcttatttc tgaggataag ctctttaaag acaaagcttt 1920
attttcatct ctcatctttt gtcctcctta gcacaatgca aaaaagaata gtaatatcag 1980
aacaggaagg aggaatggct tgctggggag cccatccagg acactgggag cacatagaga 2040
ttcacccatg tttgttgaac ttagagtcat tctcatgctt ttctttataa ttcacacata 2100
tatgcagaga agatatgttc ttgttaacat tgtatacaac atagccccaa atatagtaag 2160
atctatacta gataatccta gatgaaatgt tagagatgct atatgataca actgtggcca 2220
tgactgagga aaggagctca cgcccagaga ctgggctgct ctcccggagg ccaaacccaa 2280
gaaggtctgg caaagtcagg ctcagggaga ctctgccctg ctgcagacct cggtgtggac 2340
acacgctgca tagagctctc cttgaaaaca gaggggtctc aagacattct gcctacctat 2400
tagcttttct ttattttttt aactttttgg ggggaaaagt attttttgaga agtttgtctt 2460
gcaatgtatt tataaatagt aaataaagtt tttaccatt              2499
```

<210> 91
<211> 291
<212> PRT
<213> Homo sapiens

<400> 91

Met	Gln	Arg	Ala	Arg	Pro	Thr	Leu	Trp	Ala	Ala	Ala	Leu	Thr	Leu	Leu
1				5				10				15			

Val	Leu	Leu	Arg	Gly	Pro	Pro	Val	Ala	Arg	Ala	Gly	Ala	Ser	Ser	Gly
			20				25				30				

Gly	Leu	Gly	Pro	Val	Val	Arg	Cys	Glu	Pro	Cys	Asp	Ala	Arg	Ala	Leu
		35				40				45					

Ala	Gln	Cys	Ala	Pro	Pro	Pro	Ala	Val	Cys	Ala	Glu	Leu	Val	Arg	Glu
	50				55				60						

Pro	Gly	Cys	Gly	Cys	Cys	Leu	Thr	Cys	Ala	Leu	Ser	Glu	Gly	Gln	Pro
65				70				75				80			

Cys	Gly	Ile	Tyr	Thr	Glu	Arg	Cys	Gly	Ser	Gly	Leu	Arg	Cys	Gln	Pro

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                    85                   90                   95
Ser Pro Asp Glu Ala Arg Pro Leu Gln Ala Leu Leu Asp Gly Arg Gly
            100                 105                 110
Leu Cys Val Asn Ala Ser Ala Val Ser Arg Leu Arg Ala Tyr Leu Leu
            115                 120                 125
Pro Ala Pro Pro Ala Pro Gly Asn Ala Ser Glu Ser Glu Glu Asp Arg
        130                 135                 140
Ser Ala Gly Ser Val Glu Ser Pro Ser Val Ser Ser Thr His Arg Val
145                 150                 155                 160
Ser Asp Pro Lys Phe His Pro Leu His Ser Lys Ile Ile Ile Ile Lys
                165                 170                 175
Lys Gly His Ala Lys Asp Ser Gln Arg Tyr Lys Val Asp Tyr Glu Ser
            180                 185                 190
Gln Ser Thr Asp Thr Gln Asn Phe Ser Ser Glu Ser Lys Arg Glu Thr
        195                 200                 205
Glu Tyr Gly Pro Cys Arg Arg Glu Met Glu Asp Thr Leu Asn His Leu
        210                 215                 220
Lys Phe Leu Asn Val Leu Ser Pro Arg Gly Val His Ile Pro Asn Cys
225                 230                 235                 240
Asp Lys Lys Gly Phe Tyr Lys Lys Lys Gln Cys Arg Pro Ser Lys Gly
                245                 250                 255
Arg Lys Arg Gly Phe Cys Trp Cys Val Asp Lys Tyr Gly Gln Pro Leu
            260                 265                 270
Pro Gly Tyr Thr Thr Lys Gly Lys Glu Asp Val His Cys Tyr Ser Met
        275                 280                 285
Gln Ser Lys
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<210> 92
<211> 1639
<212> DNA
<213> Homo sapiens

<400> 92
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agagtggacc acactgcgcc aacacagaaa ttattgtaaa gctttctgat ggaagagagc 300
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gaaacttcaa gcaaatctac ttcaacactt catgtattgt gtgggtctgt tgtagggttg 480
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caaaaaacaa caaataattt ttaaatataa ggattttcct agatattgca cgggagaata 660
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gaatgggttt gctagaatgt gatatttgaa gcatcacata aaaatgatgg gacaataaat 780
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acaattgggt acccagttaa attttcattt cagataaaca acaaataatt ttttagtata 1260
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agtcttgtca ttgccagctg tgttggtagt gctgtgttga attacggaat aatgagttag 1380
aactattaaa acagccaaaa ctccacagtc aatattagta atttcttgct ggttgaaact 1440
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152

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tgtttattat gtacaaatag attcttataa tattatttaa atgactgcat ttttaaatac 1500
aaggctttat attttttaact ttaagatgtt tttatgtgct ctccaaatttt tttttactgt 1560
ttctgattgt atggaaatat aaaagtaaat atgaaacatt taaaatataa tttgttgtca 1620
aagtaaaaaa aaaaaaaaa                                               1639
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<210> 93
<211> 99
<212> PRT
<213> Homo sapiens

<400> 93
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1               5                   10                  15
Ala Ala Leu Cys Glu Gly Ala Val Leu Pro Arg Ser Ala Lys Glu Leu
            20                  25                  30
Arg Cys Gln Cys Ile Lys Thr Tyr Ser Lys Pro Phe His Pro Lys Phe
        35                  40                  45
Ile Lys Glu Leu Arg Val Ile Glu Ser Gly Pro His Cys Ala Asn Thr
    50                  55                  60
Glu Ile Ile Val Lys Leu Ser Asp Gly Arg Glu Leu Cys Leu Asp Pro
65                  70                  75                  80
Lys Glu Asn Trp Val Gln Arg Val Val Glu Lys Phe Leu Lys Arg Ala
                85                  90                  95
Glu Asn Ser
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<210> 94
<211> 1840
<212> DNA
<213> Homo sapiens

<400> 94
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aagttgctgg attatagtga ggagttcccc cacccccagga tccgaggggc acagcgcggc 180
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aatgaatgaa cttatggagc agacctcgga gatcatcacg tttgccgagt caggaacagc 480
caggaagacg ctgcacttcg agatttccaa ggaaggcagt gacctgtcag tggtggagcg 540
tgcagaagtc tggctcttcc taaaagtccc caaggccaac aggaccagga ccaaagtcac 600
catccgcctc ttccagcagc agaagcaccc gcagggcagc ttggacacag gggaagaggc 660
cgaggaagtg ggcttaaagg gggagaggag tgaactgttg ctctctgaaa aagtagtaga 720
cgctcggaag agcacctggc atgtcttccc tgtctccagc agcatccagc ggttgctgga 780
ccagggcaag agctccctgg acgttcggat tgcctgtgag cagtgccagg agagtggcgc 840
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gtttctgagt taaccagaaa aatagaaatt aaaaacaaaa caaaacaaaa aaaaaaacaa 1500
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atgtggaaat cttagcctgc cttagccagg gctcagagat gaagcagtga agagacagat 1620
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agttgtttaa acggggtatt gtcctttccc cccttgaggt tcccttgtga gcttgaatca 1740
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<210> 95
<211> 426
<212> PRT
<213> Homo sapiens

<400> 95

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Ile	Val	Arg	Ser	Ser	Pro	Thr	Pro	Gly	Ser	Glu	Gly	His	Ser	Ala	Ala
			20					25					30		
Pro	Asp	Cys	Pro	Ser	Cys	Ala	Leu	Ala	Ala	Leu	Pro	Lys	Asp	Val	Pro
		35				40					45				
Asn	Ser	Gln	Pro	Glu	Met	Val	Glu	Ala	Val	Lys	Lys	His	Ile	Leu	Asn
	50				55					60					
Met	Leu	His	Leu	Lys	Lys	Arg	Pro	Asp	Val	Thr	Gln	Pro	Val	Pro	Lys
65				70					75					80	
Ala	Ala	Leu	Leu	Asn	Ala	Ile	Arg	Lys	Leu	His	Val	Gly	Lys	Val	Gly
				85					90					95	
Glu	Asn	Gly	Tyr	Val	Glu	Ile	Glu	Asp	Asp	Ile	Gly	Arg	Arg	Ala	Glu
			100					105					110		
Met	Asn	Glu	Leu	Met	Glu	Gln	Thr	Ser	Glu	Ile	Ile	Thr	Phe	Ala	Glu
		115					120					125			
Ser	Gly	Thr	Ala	Arg	Lys	Thr	Leu	His	Phe	Glu	Ile	Ser	Lys	Glu	Gly
	130					135					140				
Ser	Asp	Leu	Ser	Val	Val	Glu	Arg	Ala	Glu	Val	Trp	Leu	Phe	Leu	Lys
145					150					155					160
Val	Pro	Lys	Ala	Asn	Arg	Thr	Arg	Thr	Lys	Val	Thr	Ile	Arg	Leu	Phe
				165					170					175	
Gln	Gln	Gln	Lys	His	Pro	Gln	Gly	Ser	Leu	Asp	Thr	Gly	Glu	Glu	Ala
			180					185					190		
Glu	Glu	Val	Gly	Leu	Lys	Gly	Glu	Arg	Ser	Glu	Leu	Leu	Leu	Ser	Glu
		195					200					205			
Lys	Val	Val	Asp	Ala	Arg	Lys	Ser	Thr	Trp	His	Val	Phe	Pro	Val	Ser
	210					215					220				
Ser	Ser	Ile	Gln	Arg	Leu	Leu	Asp	Gln	Gly	Lys	Ser	Ser	Leu	Asp	Val
225					230					235					240
Arg	Ile	Ala	Cys	Glu	Gln	Cys	Gln	Glu	Ser	Gly	Ala	Ser	Leu	Val	Leu
			245					250					255		
Leu	Gly	Lys	Lys	Lys	Lys	Lys	Glu	Glu	Glu	Gly	Glu	Gly	Lys	Lys	Lys
			260					265					270		
Gly	Gly	Gly	Glu	Gly	Gly	Ala	Gly	Ala	Asp	Glu	Glu	Lys	Glu	Gln	Ser
		275					280					285			
His	Arg	Pro	Phe	Leu	Met	Leu	Gln	Ala	Arg	Gln	Ser	Glu	Asp	His	Pro
	290					295					300				
His	Arg	Arg	Arg	Arg	Arg	Gly	Leu	Glu	Cys	Asp	Gly	Lys	Val	Asn	Ile
305					310					315					320
Cys	Cys	Lys	Lys	Gln	Phe	Phe	Val	Ser	Phe	Lys	Asp	Ile	Gly	Trp	Asn
			325					330					335		
Asp	Trp	Ile	Ile	Ala	Pro	Ser	Gly	Tyr	His	Ala	Asn	Tyr	Cys	Glu	Gly
			340					345					350		
Glu	Cys	Pro	Ser	His	Ile	Ala	Gly	Thr	Ser	Gly	Ser	Ser	Leu	Ser	Phe
		355					360					365			
His	Ser	Thr	Val	Ile	Asn	His	Tyr	Arg	Met	Arg	Gly	His	Ser	Pro	Phe

```
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Ala Asn Leu Lys Ser Cys Cys Val Pro Thr Lys Leu Arg Pro Met Ser
385                 390                    395                    400
Met Leu Tyr Tyr Asp Asp Gly Gln Asn Ile Ile Lys Lys Asp Ile Gln
                405                    410                    415
Asn Met Ile Val Glu Glu Cys Gly Cys Ser
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<210> 96
<211> 4637
<212> DNA
<213> Homo sapiens

<400> 96
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<210> 97
<211> 1051
<212> PRT
<213> Homo sapiens

<400> 97

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            20                  25                  30
Phe Asn Leu Asp Thr Arg Phe Leu Val Val Lys Glu Ala Gly Asn Pro
        35                  40                  45
Gly Ser Leu Phe Gly Tyr Ser Val Ala Leu His Arg Gln Thr Glu Arg
    50                  55                  60
Gln Gln Arg Tyr Leu Leu Leu Ala Gly Ala Pro Arg Glu Leu Ala Val
65                  70                  75                  80
Pro Asp Gly Tyr Thr Asn Arg Thr Gly Ala Val Tyr Leu Cys Pro Leu
                85                  90                  95
Thr Ala His Lys Asp Asp Cys Glu Arg Met Asn Ile Thr Val Lys Asn
            100                 105                 110
Asp Pro Gly His His Ile Ile Glu Asp Met Trp Leu Gly Val Thr Val
        115                 120                 125
Ala Ser Gln Gly Pro Ala Gly Arg Val Leu Val Cys Ala His Arg Tyr
    130                 135                 140
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Thr Gln Val Leu Trp Ser Gly Ser Glu Asp Gln Arg Arg Met Val Gly
145   150   155   160
Lys Cys Tyr Val Arg Gly Asn Asp Leu Glu Leu Asp Ser Ser Asp Asp
              165   170   175
Trp Gln Thr Tyr His Asn Glu Met Cys Asn Ser Asn Thr Asp Tyr Leu
              180   185   190
Glu Thr Gly Met Cys Gln Leu Gly Thr Ser Gly Gly Phe Thr Gln Asn
          195   200   205
Thr Val Tyr Phe Gly Ala Pro Gly Ala Tyr Asn Trp Lys Gly Asn Ser
      210   215   220
Tyr Met Ile Gln Arg Lys Glu Trp Asp Leu Ser Glu Tyr Ser Tyr Lys
225   230   235   240
Asp Pro Glu Asp Gln Gly Asn Leu Tyr Ile Gly Tyr Thr Met Gln Val
              245   250   255
Gly Ser Phe Ile Leu His Pro Lys Asn Ile Thr Ile Val Thr Gly Ala
              260   265   270
Pro Arg His Arg His Met Gly Ala Val Phe Leu Leu Ser Gln Glu Ala
      275   280   285
Gly Gly Asp Leu Arg Arg Arg Gln Val Leu Glu Gly Ser Gln Val Gly
290   295   300
Ala Tyr Phe Gly Ser Ala Ile Ala Leu Ala Asp Leu Asn Asn Asp Gly
305   310   315   320
Trp Gln Asp Leu Leu Val Gly Ala Pro Tyr Tyr Phe Glu Arg Lys Glu
              325   330   335
Glu Val Gly Gly Ala Ile Tyr Val Phe Met Asn Gln Ala Gly Thr Ser
          340   345   350
Phe Pro Ala His Pro Ser Leu Leu Leu His Gly Pro Ser Gly Ser Ala
      355   360   365
Phe Gly Leu Ser Val Ala Ser Ile Gly Asp Ile Asn Gln Asp Gly Phe
      370   375   380
Gln Asp Ile Ala Val Gly Ala Pro Phe Glu Gly Leu Gly Lys Val Tyr
385   390   395   400
Ile Tyr His Ser Ser Ser Lys Gly Leu Leu Arg Gln Pro Gln Gln Val
              405   410   415
Ile His Gly Glu Lys Leu Gly Leu Pro Gly Leu Ala Thr Phe Gly Tyr
          420   425   430
Ser Leu Ser Gly Gln Met Asp Val Asp Glu Asn Phe Tyr Pro Asp Leu
      435   440   445
Leu Val Gly Ser Leu Ser Asp His Ile Val Leu Leu Arg Ala Arg Pro
      450   455   460
Val Ile Asn Ile Val His Lys Thr Leu Val Pro Arg Pro Ala Val Leu
465   470   475   480
Asp Pro Ala Leu Cys Thr Ala Thr Ser Cys Val Gln Val Glu Leu Cys
              485   490   495
Phe Ala Tyr Asn Gln Ser Ala Gly Asn Pro Asn Tyr Arg Arg Asn Ile
              500   505   510
Thr Leu Ala Tyr Thr Leu Glu Ala Asp Arg Asp Arg Arg Pro Pro Arg
          515   520   525
Leu Arg Phe Ala Gly Ser Glu Ser Ala Val Phe His Gly Phe Phe Ser
      530   535   540
Met Pro Glu Met Arg Cys Gln Lys Leu Glu Leu Leu Leu Met Asp Asn
545   550   555   560
Leu Arg Asp Lys Leu Arg Pro Ile Ile Ile Ser Met Asn Tyr Ser Leu
              565   570   575
Pro Leu Arg Met Pro Asp Arg Pro Arg Leu Gly Leu Arg Ser Leu Asp
              580   585   590
Ala Tyr Pro Ile Leu Asn Gln Ala Gln Ala Leu Glu Asn His Thr Glu
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Val Gln Phe Gln Lys Glu Cys Gly Pro Asp Asn Lys Cys Glu Ser Asn

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Leu Gln Met Arg Ala Ala Phe Val Ser Glu Gln Gln Gln Lys Leu Ser
625                    630                    635                    640
Arg Leu Gln Tyr Ser Arg Asp Val Arg Lys Leu Leu Leu Ser Ile Asn
                 645                    650                    655
Val Thr Asn Thr Arg Thr Ser Glu Arg Ser Gly Glu Asp Ala His Glu
             660                    665                    670
Ala Leu Leu Thr Leu Val Val Pro Pro Ala Leu Leu Leu Ser Ser Val
             675                    680                    685
Arg Pro Pro Gly Ala Cys Gln Ala Asn Glu Thr Ile Phe Cys Glu Leu
         690                    695                    700
Gly Asn Pro Phe Lys Arg Asn Gln Arg Met Glu Leu Leu Ile Ala Phe
705                    710                    715                    720
Glu Val Ile Gly Val Thr Leu His Thr Arg Asp Leu Gln Val Gln Leu
                 725                    730                    735
Gln Leu Ser Thr Ser Ser His Gln Asp Asn Leu Trp Pro Met Ile Leu
             740                    745                    750
Thr Leu Leu Val Asp Tyr Thr Leu Gln Thr Ser Leu Ser Met Val Asn
             755                    760                    765
His Arg Leu Gln Ser Phe Phe Gly Gly Thr Val Met Gly Glu Ser Gly
         770                    775                    780
Met Lys Thr Val Glu Asp Val Gly Ser Pro Leu Lys Tyr Glu Phe Gln
785                    790                    795                    800
Val Gly Pro Met Gly Glu Gly Leu Val Gly Leu Gly Thr Leu Val Leu
                 805                    810                    815
Gly Leu Glu Trp Pro Tyr Glu Val Ser Asn Gly Lys Trp Leu Leu Tyr
         820                    825                    830
Pro Thr Glu Ile Thr Val His Gly Asn Gly Ser Trp Pro Cys Arg Pro
         835                    840                    845
Pro Gly Asp Leu Ile Asn Pro Leu Asn Leu Thr Leu Ser Asp Pro Gly
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Gly Gly Gln Gly Pro Pro Val Thr Leu Ala Ala Ala Lys Lys Ala
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Lys Ser Glu Thr Val Leu Thr Cys Ala Thr Gly Arg Ala His Cys Val
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Trp Leu Glu Cys Pro Ile Pro Asp Ala Pro Val Val Thr Asn Val Thr
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Val Lys Ala Arg Val Trp Asn Ser Thr Phe Ile Glu Asp Tyr Arg Asp
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Phe Asp Arg Val Arg Val Asn Gly Trp Ala Thr Leu Phe Leu Arg Thr
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Ser Ile Pro Thr Ile Asn Met Glu Asn Lys Thr Thr Trp Phe Ser Val
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Asp Ile Asp Ser Glu Leu Val Glu Glu Leu Pro Ala Glu Ile Glu Leu
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Trp Leu Val Leu Val Ala Val Gly Ala Gly Leu Leu Leu Leu Gly Leu
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<211> 4495

158

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<213> Homo sapiens

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<212> PRT
<213> Homo sapiens

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        35                  40                  45
Gly Ser Leu Phe Gly Tyr Ser Val Ala Leu His Arg Gln Thr Glu Arg
    50                  55                  60
Gln Gln Arg Tyr Leu Leu Leu Ala Gly Ala Pro Arg Glu Leu Ala Val
65                  70                  75                  80
Pro Asp Gly Tyr Thr Asn Arg Thr Gly Ala Val Tyr Leu Cys Pro Leu
                85                  90                  95
Thr Ala His Lys Asp Asp Cys Glu Arg Met Asn Ile Thr Val Lys Asn
            100                 105                 ·110
Asp Pro Gly His His Ile Ile Glu Asp.Met Trp Leu Gly Val Thr Val
        115                 120                 125
Ala Ser Gln Gly Pro Ala Gly Arg Val Leu Val Cys Ala His Arg Tyr
    130                 135                 140
Thr Gln Val Leu Trp Ser Gly Ser Glu Asp Gln Arg Arg Met Val Gly
145                 150                 155                 160
Lys Cys Tyr Val Arg Gly Asn Asp Leu Glu Leu Asp Ser Ser Asp Asp
                165                 170                 175
Trp Gln Thr Tyr His Asn Glu Met Cys Asn Ser Asn Thr Asp Tyr Leu
            180                 185                 190
Glu Thr Gly Met Cys Gln Leu Gly Thr Ser Gly Gly Phe Thr Gln Asn
        195                 200                 205
Thr Val Tyr Phe Gly Ala Pro Gly Ala Tyr Asn Trp Lys Gly Asn Ser
    210                 215                 220
Tyr Met Ile Gln Arg Lys Glu Trp Asp Leu Ser Glu Tyr Ser Tyr Lys
225                 230                 235                 240
Asp Pro Glu Asp Gln Gly Asn Leu Tyr Ile Gly Tyr Thr Met Gln Val
                245                 250                 255
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Gly Ser Phe Ile Leu His Pro Lys Asn Ile Thr Ile Val Thr Gly Ala
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Pro Arg His Arg His Met Gly Ala Val Phe Leu Leu Ser Gln Glu Ala
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Gly Gly Asp Leu Arg Arg Arg Gln Val Leu Glu Gly Ser Gln Val Gly
        290             295             300
Ala Tyr Phe Gly Ser Ala Ile Ala Leu Ala Asp Leu Asn Asn Asp Gly
305             310             315             320
Trp Gln Asp Leu Leu Val Gly Ala Pro Tyr Tyr Phe Glu Arg Lys Glu
            325             330             335
Glu Val Gly Gly Ala Ile Tyr Val Phe Met Asn Gln Ala Gly Thr Ser
            340             345             350
Phe Pro Ala His Pro Ser Leu Leu Leu His Gly Pro Ser Gly Ser Ala
            355             360             365
Phe Gly Leu Ser Val Ala Ser Ile Gly Asp Ile Asn Gln Asp Gly Phe
            370             375             380
Gln Asp Ile Ala Val Gly Ala Pro Phe Glu Gly Leu Gly Lys Val Tyr
385             390             395             400
Ile Tyr His Ser Ser Ser Lys Gly Leu Leu Arg Gln Pro Gln Gln Val
            405             410             415
Ile His Gly Glu Lys Leu Gly Leu Pro Gly Leu Ala Thr Phe Gly Tyr
            420             425             430
Ser Leu Ser Gly Gln Met Asp Val Asp Glu Asn Phe Tyr Pro Asp Leu
            435             440             445
Leu Val Gly Ser Leu Ser Asp His Ile Val Leu Leu Arg Ala Arg Pro
            450             455             460
Val Ile Asn Ile Val His Lys Thr Leu Val Pro Arg Pro Ala Val Leu
465             470             475             480
Asp Pro Ala Leu Cys Thr Ala Thr Ser Cys Val Gln Val Glu Leu Cys
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Phe Ala Tyr Asn Gln Ser Ala Gly Asn Pro Asn Tyr Arg Arg Asn Ile
            500             505             510
Thr Leu Ala Tyr Thr Leu Glu Ala Asp Arg Asp Arg Arg Pro Pro Arg
            515             520             525
Leu Arg Phe Ala Gly Ser Glu Ser Ala Val Phe His Gly Phe Phe Ser
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Met Pro Glu Met Arg Cys Gln Lys Leu Glu Leu Leu Leu Met Asp Asn
545             550             555             560
Leu Arg Asp Lys Leu Arg Pro Ile Ile Ile Ser Met Asn Tyr Ser Leu
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Pro Leu Arg Met Pro Asp Arg Pro Arg Leu Gly Leu Arg Ser Leu Asp
            580             585             590
Ala Tyr Pro Ile Leu Asn Gln Ala Gln Ala Leu Glu Asn His Thr Glu
            595             600             605
Val Gln Phe Gln Lys Glu Cys Gly Pro Asp Asn Lys Cys Glu Ser Asn
            610             615             620
Leu Gln Met Arg Ala Ala Phe Val Ser Glu Gln Gln Lys Leu Ser
625             630             635             640
Arg Leu Gln Tyr Ser Arg Asp Val Arg Lys Leu Leu Leu Ser Ile Asn
            645             650             655
Val Thr Asn Thr Arg Thr Ser Glu Arg Ser Gly Glu Asp Ala His Glu
            660             665             670
Ala Leu Leu Thr Leu Val Val Pro Pro Ala Leu Leu Leu Ser Ser Val
            675             680             685
Arg Pro Pro Gly Ala Cys Gln Ala Asn Glu Thr Ile Phe Cys Glu Leu
            690             695             700
Gly Asn Pro Phe Lys Arg Asn Gln Arg Met Glu Leu Leu Ile Ala Phe
705             710             715             720
Glu Val Ile Gly Val Thr Leu His Thr Arg Asp Leu Gln Val Gln Leu
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                     725                 730                 735
Gln Leu Ser Thr Ser Ser His Gln Asp Asn Leu Trp Pro Met Ile Leu
            740                 745                 750
Thr Leu Leu Val Asp Tyr Thr Leu Gln Thr Ser Leu Ser Met Val Asn
            755                 760                 765
His Arg Leu Gln Ser Phe Phe Gly Gly Thr Val Met Gly Glu Ser Gly
        770                 775                 780
Met Lys Thr Val Glu Asp Val Gly Ser Pro Leu Lys Tyr Glu Phe Gln
785                 790                 795                 800
Val Gly Pro Met Gly Glu Gly Leu Val Gly Leu Gly Thr Leu Val Leu
                805                 810                 815
Gly Leu Glu Trp Pro Tyr Glu Val Ser Asn Gly Lys Trp Leu Leu Tyr
            820                 825                 830
Pro Thr Glu Ile Thr Val His Gly Asn Gly Ser Trp Pro Cys Arg Pro
            835                 840                 845
Pro Gly Asp Leu Ile Asn Pro Leu Asn Leu Thr Leu Ser Asp Pro Gly
    850                 855                 860
Asp Arg Pro Ser Ser Pro Gln Arg Arg Arg Gln Leu Asp Pro Gly
865                 870                 875                 880
Gly Gly Gln Gly Pro Pro Pro Val Thr Leu Ala Ala Ala Lys Lys Ala
                885                 890                 895
Lys Ser Glu Thr Val Leu Thr Cys Ala Thr Gly Arg Ala His Cys Val
            900                 905                 910
Trp Leu Glu Cys Pro Ile Pro Asp Ala Pro Val Val Thr Asn Val Thr
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Val Lys Ala Arg Val Trp Asn Ser Thr Phe Ile Glu Asp Tyr Arg Asp
930                 935                 940
Phe Asp Arg Val Arg Val Asn Gly Trp Ala Thr Leu Phe Leu Arg Thr
945                 950                 955                 960
Ser Ile Pro Thr Ile Asn Met Glu Asn Lys Thr Thr Trp Phe Ser Val
            965                 970                 975
Asp Ile Asp Ser Glu Leu Val Glu Glu Leu Pro Ala Glu Ile Glu Leu
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Trp Leu Val Leu Val Ala Val Gly Ala Gly Leu Leu Leu Leu Gly Leu
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Ile Ile Leu Leu Leu Trp Lys Cys Asp Phe Phe Lys Arg Thr Arg Tyr
    1010                1015                1020
Tyr Gln Ile Met Pro Lys Tyr His Ala Val Arg Ile Arg Glu Glu Glu
1025                1030                1035                1040
Arg Tyr Pro Pro Pro Gly Ser Thr Leu Pro Thr Lys Lys His Trp Val
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Thr Ser Trp Gln Thr Arg Asp Gln Tyr Tyr
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<211> 4647
<212> DNA
<213> Homo sapiens

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162

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atgagatatt gtactttgt gcaatgttta ttacctttga ttaaattgta atgtgaagct 4020
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tttactaggt gaatagttca ttatgtagtg gaggcttcgt ggttgtccat tgaattgtca 4080
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attattggaa tgggatttta gaaaatagaa gttacctttg tgtgtgtttt agggaaggta 4440
gagaagaatc tgctctttct ctgaatactg ttttgacccc aggcaggacc ttggaaaggc 4500
caaaacatta acagtagtac ttctgttcac tgaagagtta tgttacatga agataaaatg 4560
gttttgtcgt gtttattatt gtattttgtg ttgatataaa taaacatggt aatttaaaca 4620
atgaaaaaaa aaaaaaaaaa aaaaaaa                                     4647
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<210> 101
<211> 788
<212> PRT
<213> Homo sapiens

<400> 101

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Met Gly Ile Glu Leu Leu Cys Leu Phe Phe Leu Phe Leu Gly Arg Asn
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Asp His Val Gln Gly Gly Cys Ala Leu Gly Gly Ala Glu Thr Cys Glu
            20                  25                  30
Asp Cys Leu Leu Ile Gly Pro Gln Cys Ala Trp Cys Ala Gln Glu Asn
        35                  40                  45
Phe Thr His Pro Ser Gly Val Gly Glu Arg Cys Asp Thr Pro Ala Asn
    50                  55                  60
Leu Leu Ala Lys Gly Cys Gln Leu Asn Phe Ile Glu Asn Pro Val Ser
65                  70                  75                  80
Gln Val Glu Ile Leu Lys Asn Lys Pro Leu Ser Val Gly Arg Gln Lys
                85                  90                  95
Asn Ser Ser Asp Ile Val Gln Ile Ala Pro Gln Ser Leu Ile Leu Lys
            100                 105                 110
Leu Arg Pro Gly Gly Ala Gln Thr Leu Gln Val His Val Arg Gln Thr
        115                 120                 125
Glu Asp Tyr Pro Val Asp Leu Tyr Tyr Leu Met Asp Leu Ser Ala Ser
        130                 135                 140
Met Asp Asp Asp Leu Asn Thr Ile Lys Glu Leu Gly Ser Arg Leu Ser
145                 150                 155                 160
Lys Glu Met Ser Lys Leu Thr Ser Asn Phe Arg Leu Gly Phe Gly Ser
                165                 170                 175
Phe Val Glu Lys Pro Val Ser Pro Phe Val Lys Thr Thr Pro Glu Glu
            180                 185                 190
Ile Ala Asn Pro Cys Ser Ser Ile Pro Tyr Phe Cys Leu Pro Thr Phe
            195                 200                 205
Gly Phe Lys His Ile Leu Pro Leu Thr Asn Asp Ala Glu Arg Phe Asn
        210                 215                 220
Glu Ile Val Lys Asn Gln Lys Ile Ser Ala Asn Ile Asp Thr Pro Glu
225                 230                 235                 240
Gly Gly Phe Asp Ala Ile Met Gln Ala Ala Val Cys Lys Glu Lys Ile
            245                 250                 255
Gly Trp Arg Asn Asp Ser Leu His Leu Leu Val Phe Val Ser Asp Ala
        260                 265                 270
Asp Ser His Phe Gly Met Asp Ser Lys Leu Ala Gly Ile Val Ile Pro
        275                 280                 285
Asn Asp Gly Leu Cys His Leu Asp Ser Lys Asn Glu Tyr Ser Met Ser
        290                 295                 300
Thr Val Leu Glu Tyr Pro Thr Ile Gly Gln Leu Ile Asp Lys Leu Val
305                 310                 315                 320
Gln Asn Asn Val Leu Leu Ile Phe Ala Val Thr Gln Glu Gln Val His
```

```
                  325                    330                    335
Leu Tyr Glu Asn Tyr Ala Lys Leu Ile Pro Gly Ala Thr Val Gly Leu
            340                    345                    350
Leu Gln Lys Asp Ser Gly Asn Ile Leu Gln Leu Ile Ile Ser Ala Tyr
            355                    360                    365
Glu Glu Leu Arg Ser Glu Val Glu Leu Glu Val Leu Gly Asp Thr Glu
        370                    375                    380
Gly Leu Asn Leu Ser Phe Thr Ala Ile Cys Asn Asn Gly Thr Leu Phe
385                    390                    395                    400
Gln His Gln Lys Lys Cys Ser His Met Lys Val Gly Asp Thr Ala Ser
                405                    410                    415
Phe Ser Val Thr Val Asn Ile Pro His Cys Glu Arg Arg Ser Arg His
            420                    425                    430
Ile Ile Ile Lys Pro Val Gly Leu Gly Asp Ala Leu Glu Leu Leu Val
            435                    440                    445
Ser Pro Glu Cys Asn Cys Asp Cys Gln Lys Glu Val Glu Val Asn Ser
        450                    455                    460
Ser Lys Cys His His Gly Asn Gly Ser Phe Gln Cys Gly Val Cys Ala
465                    470                    475                    480
Cys His Pro Gly His Met Gly Pro Arg Cys Glu Cys Gly Glu Asp Met
                485                    490                    495
Leu Ser Thr Asp Ser Cys Lys Glu Ala Pro Asp His Pro Ser Cys Ser
            500                    505                    510
Gly Arg Gly Asp Cys Tyr Cys Gly Gln Cys Ile Cys His Leu Ser Pro
            515                    520                    525
Tyr Gly Asn Ile Tyr Gly Pro Tyr Cys Gln Cys Asp Asn Phe Ser Cys
        530                    535                    540
Val Arg His Lys Gly Leu Leu Cys Gly Gly Asn Gly Asp Cys Asp Cys
545                    550                    555                    560
Gly Glu Cys Val Cys Arg Ser Gly Trp Thr Gly Glu Tyr Cys Asn Cys
                565                    570                    575
Thr Thr Ser Thr Asp Ser Cys Val Ser Glu Asp Gly Val Leu Cys Ser
            580                    585                    590
Gly Arg Gly Asp Cys Val Cys Gly Lys Cys Val Cys Thr Asn Pro Gly
        595                    600                    605
Ala Ser Gly Pro Thr Cys Glu Arg Cys Pro Thr Cys Gly Asp Pro Cys
610                    615                    620
Asn Ser Lys Arg Ser Cys Ile Glu Cys His Leu Ser Ala Ala Gly Gln
625                    630                    635                    640
Ala Arg Glu Glu Cys Val Asp Lys Cys Lys Leu Ala Gly Ala Thr Ile
                645                    650                    655
Ser Glu Glu Glu Asp Phe Ser Lys Asp Gly Ser Val Ser Cys Ser Leu
            660                    665                    670
Gln Gly Glu Asn Glu Cys Leu Ile Thr Phe Leu Ile Thr Thr Asp Asn
            675                    680                    685
Glu Gly Lys Thr Ile Ile His Ser Ile Asn Glu Lys Asp Cys Pro Lys
        690                    695                    700
Pro Pro Asn Ile Pro Met Ile Met Leu Gly Val Ser Leu Ala Ile Leu
705                    710                    715                    720
Leu Ile Gly Val Val Leu Leu Cys Ile Trp Lys Leu Leu Val Ser Phe
                725                    730                    735
His Asp Arg Lys Glu Val Ala Lys Phe Glu Ala Glu Arg Ser Lys Ala
            740                    745                    750
Lys Trp Gln Thr Gly Thr Asn Pro Leu Tyr Arg Gly Ser Thr Ser Thr
            755                    760                    765
Phe Lys Asn Val Thr Tyr Lys His Arg Glu Lys Gln Lys Val Asp Leu
770                    775                    780
Ser Thr Asp Cys
785
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```
<210> 102
<211> 2231
<212> DNA
<213> Homo sapiens

<400> 102
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gcggggcggg aggggcgcg  gcgaggcccg ccggcggggc aaaaccggcc tgggccctgg 180
cggccgcagg agcgcgtgcg gcgtggactt tgccgggctc gccacacagc cccagacccg 240
tttaggaccg ggagaccgaa cgcagcgtcc agccggggag tttcggcggc gttctccggg 300
caccgcgcgc gggaagccag acgcagcggg gggacacatc tcgcggtggc gttgccagag 360
tgaggagtta gcaggcagga cttgacgagg ctctttggtt tttctagtcc tcaaccactg 420
aagaagaagc ttgatgcttg gctgtcagaa gacatgaatt acgcacggtt catcacggca 480
gcgagcgcag ccagaaaccc ttctcccatc cggaccatga ctgacatatt gagcagagga 540
ccaaaatcga tgatctcctt ggctggtggc ttaccaaatc caaacatgtt tccttttaag 600
actgccgtaa tcactgtaga aaatggaaag accatccaat ttggagaaga gatgatgaag 660
agagcacttc agtattctcc gagtgctgga attccagagc ttttgtcctg gctaaaacag 720
ttacaaataa aattgcataa tcctcctacc atccattacc cacccagtca aggacaaatg 780
gatctatgtg tcacatctgg cagccaacaa ggtctttgta aggtgtttga aatgatcatt 840
aatcctggag ataatgtcct cctagatgaa cctgcttatt caggaactct tcaaagtctg 900
cacccactgg gctgcaacat tattaatgtt gccagtgatg aaagtgggat tgttccagat 960
tccctaagag acatactttc cagatggaaa ccagaagatg caaagaatcc ccagaaaaac 1020
accccaaat  ttctttatac tgttccaaat ggcaacaacc ctactggaaa ctcattaacc 1080
agtgaacgca aaaaggaaat ctatgagctt gcaagaaaat atgatttcct cataatagaa 1140
gatgatcctt actattttct ccagtttaac aagttcaggg taccaacatt tctttccatg 1200
gatgttgatg gacgtgtcat cagagctgac tcttttcaa  aaatcatttc ctctgggttg 1260
agaataggat ttttaactgg tccaaaaccc ttaatagaga gagttatttt acacatacaa 1320
gtttcaacat tgcacccag  cacttttaac cagctcatga tatcacagct tctacacgaa 1380
tggggagaag aaggtttcat ggctcatgta gacagggtta ttgatttcta tagtaaccag 1440
aaggatgcaa tactggcagc tgcagacaag tggttaactg gtttggcaga atggcatgtt 1500
cctgctgctg gaatgttttt atggattaaa gttaaaggca ttaatgatgt aaaagaactg 1560
attgaagaaa aggccgttaa gatgggggta ttaatgctcc ctggaaatgc tttctacgtc 1620
gatagctcag ctcctagccc ttacttgaga gcatccttct cttcagcttc tccagaacag 1680
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aggatcactt gaacccagga attcaggctg cagtaagcta cgatcacacc actgcactct 1860
ggcctgcatg cactctggcc tgcatggcag aacaagaccc tgtctctaaa aaagagaaa  1920
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gtagttcaac tagacacctt ttaaagtgcc tctaaattca tcagatggcc aaactgtatt 2100
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aaaaaaaaaa a                                                      2231

<210> 103
<211> 425
<212> PRT
<213> Homo sapiens

<400> 103
Met Asn Tyr Ala Arg Phe Ile Thr Ala Ala Ser Ala Ala Arg Asn Pro
1               5                   10                  15
Ser Pro Ile Arg Thr Met Thr Asp Ile Leu Ser Arg Gly Pro Lys Ser
            20                  25                  30
Met Ile Ser Leu Ala Gly Gly Leu Pro Asn Pro Asn Met Phe Pro Phe
        35                  40                  45
```

```
Lys Thr Ala Val Ile Thr Val Glu Asn Gly Lys Thr Ile Gln Phe Gly
    50                  55                  60
Glu Glu Met Met Lys Arg Ala Leu Gln Tyr Ser Pro Ser Ala Gly Ile
65                  70                  75                  80
Pro Glu Leu Leu Ser Trp Leu Lys Gln Leu Gln Ile Lys Leu His Asn
                85                  90                  95
Pro Pro Thr Ile His Tyr Pro Pro Ser Gln Gly Gln Met Asp Leu Cys
            100                 105                 110
Val Thr Ser Gly Ser Gln Gln Gly Leu Cys Lys Val Phe Glu Met Ile
        115                 120                 125
Ile Asn Pro Gly Asp Asn Val Leu Leu Asp Glu Pro Ala Tyr Ser Gly
    130                 135                 140
Thr Leu Gln Ser Leu His Pro Leu Gly Cys Asn Ile Ile Asn Val Ala
145                 150                 155                 160
Ser Asp Glu Ser Gly Ile Val Pro Asp Ser Leu Arg Asp Ile Leu Ser
                165                 170                 175
Arg Trp Lys Pro Glu Asp Ala Lys Asn Pro Gln Lys Asn Thr Pro Lys
            180                 185                 190
Phe Leu Tyr Thr Val Pro Asn Gly Asn Asn Pro Thr Gly Asn Ser Leu
        195                 200                 205
Thr Ser Glu Arg Lys Lys Glu Ile Tyr Glu Leu Ala Arg Lys Tyr Asp
    210                 215                 220
Phe Leu Ile Ile Glu Asp Asp Pro Tyr Tyr Phe Leu Gln Pne Asn Lys
225                 230                 235                 240
Phe Arg Val Pro Thr Phe Leu Ser Met Asp Val Asp Gly Arg Val Ile
                245                 250                 255
Arg Ala Asp Ser Phe Ser Lys Ile Ile Ser Ser Gly Leu Arg Ile Gly
            260                 265                 270
Phe Leu Thr Gly Pro Lys Pro Leu Ile Glu Arg Val Ile Leu His Ile
        275                 280                 285
Gln Val Ser Thr Leu His Pro Ser Thr Phe Asn Gln Leu Met Ile Ser
    290                 295                 300
Gln Leu Leu His Glu Trp Gly Glu Glu Gly Phe Met Ala His Val Asp
305                 310                 315                 320
Arg Val Ile Asp Phe Tyr Ser Asn Gln Lys Asp Ala Ile Leu Ala Ala
                325                 330                 335
Ala Asp Lys Trp Leu Thr Gly Leu Ala Glu Trp His Val Pro Ala Ala
            340                 345                 350
Gly Met Phe Leu Trp Ile Lys Val Lys Gly Ile Asn Asp Val Lys Glu
        355                 360                 365
Leu Ile Glu Glu Lys Ala Val Lys Met Gly Val Leu Met Leu Pro Gly
    370                 375                 380
Asn Ala Phe Tyr Val Asp Ser Ser Ala Pro Ser Pro Tyr Leu Arg Ala
385                 390                 395                 400
Ser Phe Ser Ser Ala Ser Pro Glu Gln Met Asp Val Ala Phe Gln Val
                405                 410                 415
Leu Ala Gln Leu Ile Lys Glu Ser Leu
            420                 425
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<210> 104
<211> 3176
<212> DNA
<213> Homo sapiens

<400> 104
tgataaccca aggtattcac agcaagatac agtgagtctt aaagttaagc accgtgcaat 60
tagctttgct tccttgggtt tttgaaacat gcatctgtat aaacctgcct gtgcagacat 120
cccgagcccc aagctgggtc tgccaaaatc cagtgaatcg gctctaaaat gtagatggca 180
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cctagcagtg accaagactc agcctcaggc ggcctgcaaa cctgtgaggc ccagtggagc 240
agccgaacag aaatatgtgg aaaagtttct acgtgttcat ggaatttcgt tgcaggaaac 300
caccagagca gagacgggca tggcatacag gaatcttgga aaatcaggac tcagagtttc 360
ttgcttgggt cttggaacat gggtgacatt tggaggtcaa atttcagatg aggttgctga 420
acggctgatg accatcgcct atgaaagtgg tgttaacctc tttgatactg ccgaagtcta 480
tgctgctgga aaggctgaag tgattctggg gagcatcatc aagaagaaag gctggaggag 540
gtccagtctg gtcataacaa ccaaactcta ctggggtgga aagctgaaa cagaaagagg 600
gctgtcaaga aagcatatta ttgaaggatt gaagggctcc ctccagaggc tgcagctcga 660
gtatgtggat gtggtctttg caaatcgacc ggacagtaac actcccatgg aagaaattgt 720
ccgagccatg acacatgtga taaaccaagg catggcgatg tactggggca·cctcgagatg 780
gagtgctatg gagatcatgg aagcctattc tgtagcaaga cagttcaata tgatcccacc 840
ggtctgtgaa caagctgagt accatctttt ccagagagag aaagtggagg tccagctgcc 900
agagctctac cacaaaatag gtgttggcgc aatgacatgg tctccacttg cctgtggaat 960
catctcagga aaatacggaa acggggtgcc tgaaagttcc agggcttcac tgaagtgcta 1020
ccagtggttg aaagaaagaa ttgtaagtga agaagggaga aaacagcaaa acaagctaaa 1080
agacctttcc ccaattgcgg agcgtctggg atgcacacta cctcagctag ctgttgcgtg 1140
gtgcctgaga aatgaaggtg tgagttctgt gctcctggga tcatccactc ctgaacaact 1200
cattgaaaac cttggtgcca ttcaggttct cccaaagatg acatcacatg tggtaaatga 1260
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tactgttgct tggccccaag aatggccttg tacaacttat ccagaatgtc tattaggatt 2040
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tcaatccagc tgaactgctg tatgtataga ggagcttgag gtgctgtcta atgggaaatg 3120
tgatttgatt gatttatttg cttagagtaa taaaagcatt ttgtgcattc aatctt    3176
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<210> 105
<211> 408
<212> PRT
<213> Homo sapiens

<400> 105
Met His Leu Tyr Lys Pro Ala Cys Ala Asp Ile Pro Ser Pro Lys Leu
1   5   10   15

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Gly Leu Pro Lys Ser Ser Glu Ser Ala Leu Lys Cys Arg Trp His Leu
        20              25              30
Ala Val Thr Lys Thr Gln Pro Gln Ala Ala Cys Lys Pro Val Arg Pro
        35              40              45
Ser Gly Ala Ala Glu Gln Lys Tyr Val Glu Lys Phe Leu Arg Val His
    50              55              60
Gly Ile Ser Leu Gln Glu Thr Thr Arg Ala Glu Thr Gly Met Ala Tyr
65              70              75              80
Arg Asn Leu Gly Lys Ser Gly Leu Arg Val Ser Cys Leu Gly Leu Gly
            85              90              95
Thr Trp Val Thr Phe Gly Gly Gln Ile Ser Asp Glu Val Ala Glu Arg
            100             105             110
Leu Met Thr Ile Ala Tyr Glu Ser Gly Val Asn Leu Phe Asp Thr Ala
        115             120             125
Glu Val Tyr Ala Ala Gly Lys Ala Glu Val Ile Leu Gly Ser Ile Ile
    130             135             140
Lys Lys Lys Gly Trp Arg Arg Ser Ser Leu Val Ile Thr Thr Lys Leu
145             150             155             160
Tyr Trp Gly Gly Lys Ala Glu Thr Glu Arg Gly Leu Ser Arg Lys His
            165             170             175
Ile Ile Glu Gly Leu Lys Gly Ser Leu Gln Arg Leu Gln Leu Glu Tyr
            180             185             190
Val Asp Val Val Phe Ala Asn Arg Pro Asp Ser Asn Thr Pro Met Glu
        195             200             205
Glu Ile Val Arg Ala Met Thr His Val Ile Asn Gln Gly Met Ala Met
    210             215             220
Tyr Trp Gly Thr Ser Arg Trp Ser Ala Met Glu Ile Met Glu Ala Tyr
225             230             235             240
Ser Val Ala Arg Gln Phe Asn Met Ile Pro Pro Val Cys Glu Gln Ala
            245             250             255
Glu Tyr His Leu Phe Gln Arg Glu Lys Val Glu Val Gln Leu Pro Glu
        260             265             270
Leu Tyr His Lys Ile Gly Val Gly Ala Met Thr Trp Ser Pro Leu Ala
        275             280             285
Cys Gly Ile Ile Ser Gly Lys Tyr Gly Asn Gly Val Pro Glu Ser Ser
    290             295             300
Arg Ala Ser Leu Lys Cys Tyr Gln Trp Leu Lys Glu Arg Ile Val Ser
305             310             315             320
Glu Glu Gly Arg Lys Gln Gln Asn Lys Leu Lys Asp Leu Ser Pro Ile
            325             330             335
Ala Glu Arg Leu Gly Cys Thr Leu Pro Gln Leu Ala Val Ala Trp Cys
        340             345             350
Leu Arg Asn Glu Gly Val Ser Ser Val Leu Leu Gly Ser Ser Thr Pro
        355             360             365
Glu Gln Leu Ile Glu Asn Leu Gly Ala Ile Gln Val Leu Pro Lys Met
    370             375             380
Thr Ser His Val Val Asn Glu Ile Asp Asn Ile Leu Arg Asn Lys Pro
385             390             395             400
Tyr Ser Lys Lys Asp Tyr Arg Ser
            405
```

<210> 106
<211> 3103
<212> DNA
<213> Homo sapiens

<400> 106
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agtctgggtt ttctgtagca gggaaagaca aatctcccaa gaaagcctca gaaaacgcta 180
aagacagcag ccttagtccc tcaggggaaa gccagctcag ggcgcgtcaa ctggctctgc 240
tgcgcgaagt ggagatgaac tggtacctaa agctctgcga cctgtccagc gagcacacca 300
ccgtctgcac cacaggcatg ccgcacagga atcttggaaa atcaggactc agagtttctt 360
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ccagtctggt cataacaacc aaactctact ggggtggaaa agctgaaaca gaaagagggc 600
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atgtggatgt ggtctttgca aatcgaccgg acagtaacac tcccatggaa gaaattgtcc 720
gagccatgac acatgtgata aaccaaggca tggcgatgta ctggggcacc tcgagatgga 780
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agctctacca caaaataggt gttggcgcaa tgacatggtc tccacttgcc tgtggaatca 960
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agtggttgaa agaaagaatt gtaagtgaag aagggagaaa acagcaaaac aagctaaaag 1080
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catgaaccac agaagctgca tggttaaaat agcggcctgt gcccagtaca gaaaggtgtt 1380
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agaattttat atatttttt tctatttca cattcatact aacaagtttt gttccatttg 1920
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tgcttggccc caagaatggc cttgtacaac ttatccagaa tgtctattag gattctaatg 2040
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cagctgattt atgaaagaga gggctacact gctatggaaa cttagcttca aagaaaatgc 2160
aatgtatctg caattaggtg ttcatttttt actacatttt attaaaacct gctttatact 2220
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gttatcagaa tgatggaaat tgatcatttt cagttgttca ttgtgtattc aatccagcga 3060
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<210> 107
<211> 419
<212> PRT
<213> Homo sapiens

<400> 107
Met Leu Ala Ala Arg Thr Gly Ala Ala Gly Ser Gln Ile Ser Glu Glu

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Asn Thr Lys Leu Arg Arg Gln Ser Gly Phe Ser Val Ala Gly Lys Asp
            20                    25                    30
Lys Ser Pro Lys Lys Ala Ser Glu Asn Ala Lys Asp Ser Ser Leu Ser
        35                    40                    45
Pro Ser Gly Glu Ser Gln Leu Arg Ala Arg Gln Leu Ala Leu Leu Arg
    50                    55                    60
Glu Val Glu Met Asn Trp Tyr Leu Lys Leu Cys Asp Leu Ser Ser Glu
65                    70                    75                    80
His Thr Thr Val Cys Thr Thr Gly Met Pro His Arg Asn Leu Gly Lys
            85                    90                    95
Ser Gly Leu Arg Val Ser Cys Leu Gly Leu Gly Thr Trp Val Thr Phe
            100                   105                   110
Gly Gly Gln Ile Ser Asp Glu Val Ala Glu Arg Leu Met Thr Ile Ala
        115                   120                   125
Tyr Glu Ser Gly Val Asn Leu Phe Asp Thr Ala Glu Val Tyr Ala Ala
    130                   135                   140
Gly Lys Ala Glu Val Ile Leu Gly Ser Ile Ile Lys Lys Lys Gly Trp
145                   150                   155                   160
Arg Arg Ser Ser Leu Val Ile Thr Thr Lys Leu Tyr Trp Gly Gly Lys
            165                   170                   175
Ala Glu Thr Glu Arg Gly Leu Ser Arg Lys His Ile Ile Glu Gly Leu
            180                   185                   190
Lys Gly Ser Leu Gln Arg Leu Gln Leu Glu Tyr Val Asp Val Val Phe
        195                   200                   205
Ala Asn Arg Pro Asp Ser Asn Thr Pro Met Glu Glu Ile Val Arg Ala
    210                   215                   220
Met Thr His Val Ile Asn Gln Gly Met Ala Met Tyr Trp Gly Thr Ser
225                   230                   235                   240
Arg Trp Ser Ala Met Glu Ile Met Glu Ala Tyr Ser Val Ala Arg Gln
            245                   250                   255
Phe Asn Met Ile Pro Pro Val Cys Glu Gln Ala Glu Tyr His Leu Phe
            260                   265                   270
Gln Arg Glu Lys Val Glu Val Gln Leu Pro Glu Leu Tyr His Lys Ile
        275                   280                   285
Gly Val Gly Ala Met Thr Trp Ser Pro Leu Ala Cys Gly Ile Ile Ser
    290                   295                   300
Gly Lys Tyr Gly Asn Gly Val Pro Glu Ser Ser Arg Ala Ser Leu Lys
305                   310                   315                   320
Cys Tyr Gln Trp Leu Lys Glu Arg Ile Val Ser Glu Glu Gly Arg Lys
            325                   330                   335
Gln Gln Asn Lys Leu Lys Asp Leu Ser Pro Ile Ala Glu Arg Leu Gly
            340                   345                   350
Cys Thr Leu Pro Gln Leu Ala Val Ala Trp Cys Leu Arg Asn Glu Gly
        355                   360                   365
Val Ser Ser Val Leu Leu Gly Ser Ser Thr Pro Glu Gln Leu Ile Glu
    370                   375                   380
Asn Leu Gly Ala Ile Gln Val Leu Pro Lys Met Thr Ser His Val Val
385                   390                   395                   400
Asn Glu Ile Asp Asn Ile Leu Arg Asn Lys Pro Tyr Ser Lys Lys Asp
            405                   410                   415
Tyr Arg Ser
```

<210> 108
<211> 2620
<212> DNA
<213> Homo sapiens

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<400> 108
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gtggtgaacg cagcccgggc caaattccgc acggtcgcta tcatcgcgcg cagcctgggg 180
acgttcacgc ctcagcatca catttctctc aaagagtcca ccgcaaagca gactggcatg 240
aaatatagga atcttggaaa atcaggactc agagtttctt gcttgggtct tggaacatgg 300
gtgacatttg gaggtcaaat ttcagatgag gttgctgaac ggctgatgac catcgcctat 360
gaaagtggtg ttaacctctt tgatactgcc gaagtctatg ctgctggaaa ggctgaagtg 420
attctgggga gcatcatcaa gaagaaaggc tggaggaggt ccagtctggt cataacaacc 480
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gaaggattga agggctccct ccagaggctg cagctcgagt atgtggatgt ggtctttgca 600
aatcgaccgg acagtaacac tcccatggaa gaaattgtcc gagccatgac acatgtgata 660
aaccaaggca tggcgatgta ctggggcacc tcgagatgga gtgctatgga gatcatggaa 720
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aagaatatgg gggccagggg gtgtggtact accttcaggc atttggtaac tcaaagaagg 1560
ctgtacagat atatttttc aaaaagaaca aaatccacag atgcaatgtg agttgcgtaa 1620
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attgtgagac tatactgtgt caatatctgt aaaaagagag                      2620
```

```
<210> 109
<211> 401
<212> PRT
<213> Homo sapiens
```

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<400> 109
Met Gln Val Ser Ile Ala Cys Thr Glu His Asn Leu Lys Ser Arg Asn
1               5                   10                  15
Gly Glu Asp Arg Leu Leu Ser Lys Gln Ser Ser Thr Ala Pro Asn Val
            20                  25                  30
Val Asn Ala Ala Arg Ala Lys Phe Arg Thr Val Ala Ile Ile Ala Arg
        35                  40                  45
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Ser Leu Gly Thr Phe Thr Pro Gln His His Ile Ser Leu Lys Glu Ser
    50                  55              60
Thr Ala Lys Gln Thr Gly Met Lys Tyr Arg Asn Leu Gly Lys Ser Gly
65                  70              75                  80
Leu Arg Val Ser Cys Leu Gly Leu Gly Thr Trp Val Thr Phe Gly Gly
            85                  90                  95
Gln Ile Ser Asp Glu Val Ala Glu Arg Leu Met Thr Ile Ala Tyr Glu
            100             105             110
Ser Gly Val Asn Leu Phe Asp Thr Ala Glu Val Tyr Ala Ala Gly Lys
        115             120             125
Ala Glu Val Ile Leu Gly Ser Ile Ile Lys Lys Lys Gly Trp Arg Arg
    130             135             140
Ser Ser Leu Val Ile Thr Thr Lys Leu Tyr Trp Gly Gly Lys Ala Glu
145             150             155                 160
Thr Glu Arg Gly Leu Ser Arg Lys His Ile Ile Glu Gly Leu Lys Gly
            165             170             175
Ser Leu Gln Arg Leu Gln Leu Glu Tyr Val Asp Val Val Phe Ala Asn
        180             185             190
Arg Pro Asp Ser Asn Thr Pro Met Glu Glu Ile Val Arg Ala Met Thr
        195             200             205
His Val Ile Asn Gln Gly Met Ala Met Tyr Trp Gly Thr Ser Arg Trp
    210             215             220
Ser Ala Met Glu Ile Met Glu Ala Tyr Ser Val Ala Arg Gln Phe Asn
225             230             235                 240
Met Ile Pro Pro Val Cys Glu Gln Ala Glu Tyr His Leu Phe Gln Arg
            245             250             255
Glu Lys Val Glu Val Gln Leu Pro Glu Leu Tyr His Lys Ile Gly Val
        260             265             270
Gly Ala Met Thr Trp Ser Pro Leu Ala Cys Gly Ile Ile Ser Gly Lys
    275             280             285
Tyr Gly Asn Gly Val Pro Glu Ser Ser Arg Ala Ser Leu Lys Cys Tyr
290             295             300
Gln Trp Leu Lys Glu Arg Ile Val Ser Glu Glu Gly Arg Lys Gln Gln
305             310             315                 320
Asn Lys Leu Lys Asp Leu Ser Pro Ile Ala Glu Arg Leu Gly Cys Thr
            325             330             335
Leu Pro Gln Leu Ala Val Ala Trp Cys Leu Arg Asn Glu Gly Val Ser
        340             345             350
Ser Val Leu Leu Gly Ser Ser Thr Pro Glu Gln Leu Ile Glu Asn Leu
        355             360             365
Gly Ala Ile Gln Val Leu Pro Lys Met Thr Ser His Val Val Asn Glu
370             375             380
Ile Asp Asn Ile Leu Arg Asn Lys Pro Tyr Ser Lys Lys Asp Tyr Arg
385             390             395                 400
Ser
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<210> 110
<211> 3944
<212> DNA
<213> Homo sapiens

<400> 110
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cttcaaacct tcacagctaa tcaaagacct ggccaaagag atccggctca gtgagaatgc 60
ctccaaagcc gtccgaccgg aagtgaatac tgtcgcctcg tcagatgagg tgtgtgacgg 120
ggaccgggag aaggaggagc ccccgtctcc cattgaggcc accccgcctc aatccctcct 180
ggagaaagtg tccaaaaaaa agactcccaa aactgtgaag atgcccaagc catccaaaat 240
ccccaagccc ccgaagcccc ctaagccccc aaggcccccc aaaacgctga agctcaaaga 300
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tggaggcaag aagaaaggga agaagtcccg ggagtcagcc tcacccacca tccccaacct 360
ggacctgctc gaagcccaca ccaaggaggc actgaccaag atggagccgc ccaagaaggg 420
caaggccaca aagagtgtcc tgagtgtgcc caacaaagat gtggttcaca tgcagaatga 480
tgtggagagg ctggaaattc gagagcaaac caagagcaag tcagaggcca agtggaagta 540
caagaacagc aaacctgact ccttactgaa gatggaagag gagcagaagc tagagaagtc 600
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tgttttattt cccaattcat attactcttg tatcgagtcc atgaggtcta aggcaactta 3900
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<210> 111
<211> 677
<212> PRT
<213> Homo sapiens

<400> 111
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Ser Glu Asn Ala Ser Lys Ala Val Arg Pro Glu Val Asn Thr Val Ala
            20                  25                  30
Ser Ser Asp Glu Val Cys Asp Gly Asp Arg Glu Lys Glu Glu Pro Pro
            35                  40                  45
Ser Pro Ile Glu Ala Thr Pro Pro Gln Ser Leu Leu Glu Lys Val Ser
        50                  55                  60
Lys Lys Lys Thr Pro Lys Thr Val Lys Met Pro Lys Pro Ser Lys Ile
65                  70                  75                  80
Pro Lys Pro Pro Lys Pro Pro Lys Pro Pro Arg Pro Pro Lys Thr Leu
                85                  90                  95
Lys Leu Lys Asp Gly Gly Lys Lys Lys Gly Lys Lys Ser Arg Glu Ser
            100                 105                 110
Ala Ser Pro Thr Ile Pro Asn Leu Asp Leu Leu Glu Ala His Thr Lys
            115                 120                 125
Glu Ala Leu Thr Lys Met Glu Pro Pro Lys Lys Gly Lys Ala Thr Lys
        130                 135                 140
Ser Val Leu Ser Val Pro Asn Lys Asp Val Val His Met Gln Asn Asp
145                 150                 155                 160
Val Glu Arg Leu Glu Ile Arg Glu Gln Thr Lys Ser Lys Ser Glu Ala
                165                 170                 175
Lys Trp Lys Tyr Lys Asn Ser Lys Pro Asp Ser Leu Leu Lys Met Glu
            180                 185                 190
Glu Glu Gln Lys Leu Glu Lys Ser Pro Leu Ala Gly Asn Lys Asp Asn
            195                 200                 205
Lys Phe Ser Phe Ser Phe Ser Asn Lys Lys Leu Leu Gly Ser Lys Ala
        210                 215                 220
Leu Arg Pro Pro Thr Ser Pro Gly Val Phe Gly Ala Leu Gln Asn Phe
225                 230                 235                 240
Lys Glu Asp Lys Pro Lys Pro Val Arg Asp Glu Tyr Glu Tyr Val Ser
                245                 250                 255
Asp Asp Gly Glu Leu Lys Ile Asp Glu Phe Pro Ile Arg Arg Lys Lys
                260                 265                 270
Asn Ala Pro Lys Arg Asp Leu Ser Phe Leu Leu Asp Lys Lys Ala Val
            275                 280                 285
Leu Pro Thr Pro Val Thr Lys Pro Lys Leu Asp Ser Ala Ala Tyr Lys
        290                 295                 300
Gln Ser Asp Asp Ser Ser Asp Glu Gly Ser Leu His Ile Asp Thr Asp
305                 310                 315                 320
Thr Lys Pro Gly Arg Asn Ala Arg Val Lys Lys Glu Ser Gly Ser Ser
                325                 330                 335
Ala Ala Gly Ile Leu Asp Leu Leu Gln Ala Ser Glu Glu Val Gly Ala
            340                 345                 350
Leu Glu Tyr Asn Pro Ser Ser Gln Pro Pro Ala Ser Pro Ser Thr Gln
            355                 360                 365
Glu Ala Ile Gln Gly Met Leu Ser Met Ala Asn Leu Gln Ala Ser Asp
        370                 375                 380
Ser Cys Leu Gln Thr Thr Trp Gly Ala Gly Gln Ala Lys Gly Ser Ser
385                 390                 395                 400
```

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Leu Ala Ala His Gly Ala Arg Lys Asn Gly Gly Gly Ser Gly Lys Ser
            405             410                 415
Ala Gly Lys Arg Leu Leu Lys Arg Ala Ala Lys Asn Ser Val Asp Leu
            420             425             430
Asp Asp Tyr Glu Glu Glu Gln Asp His Leu Asp Ala Cys Phe Lys Asp
            435             440             445
Ser Asp Tyr Val Tyr Pro Ser Leu Glu Ser Asp Glu Asp Asn Pro Ile
        450             455             460
Phe Lys Ser Arg Ser Lys Lys Arg Lys Gly Ser Asp Asp Ala Pro Tyr
465             470             475                 480
Ser Pro Thr Ala Arg Val Gly Pro Ser Val Pro Arg Gln Asp Arg Pro
                485             490             495
Val Arg Glu Gly Thr Arg Val Ala Ser Ile Glu Thr Gly Leu Ala Ala
            500             505             510
Ala Ala Ala Lys Leu Ser Gln Gln Glu Glu Gln Lys Ser Lys Lys Lys
        515             520             525
Lys Ser Ala Lys Arg Lys Leu Thr Pro Asn Thr Thr Ser Pro Ser Thr
    530             535             540
Ser Thr Ser Ile Ser Ala Gly Thr Thr Ser Thr Ser Thr Thr Pro Ala
545             550             555                 560
Ser Thr Thr Pro Ala Ser Thr Thr Pro Ala Ser Thr Thr Pro Ala Ser
            565             570             575
Thr Ser Thr Ala Ser Ser Gln Ala Ser Gln Glu Gly Ser Ser Pro Glu
        580             585             590
Pro Pro Pro Glu Ser His Ser Ser Ser Leu Ala Asp His Glu Tyr Thr
    595             600             605
Ala Ala Gly Thr Phe Thr Gly Ala Gln Ala Gly Arg Thr Ser Gln Pro
    610             615             620
Met Ala Pro Gly Val Phe Leu Thr Gln Arg Arg Pro Ser Ala Ser Ser
625             630             635                 640
Pro Asn Asn Asn Thr Ala Ala Lys Gly Lys Arg Thr Lys Lys Gly Met
            645             650             655
Ala Thr Ala Lys Gln Arg Leu Gly Lys Ile Leu Lys Ile His Arg Asn
            660             665             670
Gly Lys Leu Leu Leu
            675
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<210> 112
<211> 5433
<212> DNA
<213> Homo sapiens

<400> 112
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gagtttagac ccagccaggg ttgtagccct ggatactatc gggatcataa aggcttgtat 180
accggacggt gtgttccctg caattgcaac ggacattcaa atcaatgcca ggatggctca 240
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ctgactggag actgcataaa ccaagaaccc aaagatagca gccctgcaga agaatgtgat 600
gattgcgaca gctgtgtgat gaccctcctg aacgacctgg ccaccatggg cgagcagctc 660
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<213> Homo sapiens

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Ser Gln Leu Gln Ala Ser Tyr Val Glu Phe Arg Pro Ser Gln Gly Cys
        35                  40                  45
Ser Pro Gly Tyr Tyr Arg Asp His Lys Gly Leu Tyr Thr Gly Arg Cys
    50                  55                  60
Val Pro Cys Asn Cys Asn Gly His Ser Asn Gln Cys Gln Asp Gly Ser
65                  70                  75                  80
Gly Ile Cys Val Asn Cys Gln His Asn Thr Ala Gly Glu His Cys Glu
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Arg Cys Gln Glu Gly Tyr Tyr Gly Asn Ala Val His Gly Ser Cys Arg
            100                 105                 110
Ala Cys Pro Cys Pro His Thr Asn Ser Phe Ala Thr Gly Cys Val Val
        115                 120                 125
Asn Gly Gly Asp Val Arg Cys Ser Cys Lys Ala Gly Tyr Thr Gly Thr
    130                 135                 140
Gln Cys Glu Arg Cys Ala Pro Gly Tyr Phe Gly Asn Pro Gln Lys Phe
145                 150                 155                 160
Gly Gly Ser Cys Gln Pro Cys Ser Cys Asn Ser Asn Gly Gln Leu Gly
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Ser Cys His Pro Leu Thr Gly Asp Cys Ile Asn Gln Glu Pro Lys Asp
            180                 185                 190
Ser Ser Pro Ala Glu Glu Cys Asp Asp Cys Asp Ser Cys Val Met Thr
        195                 200                 205
Leu Leu Asn Asp Leu Ala Thr Met Gly Glu Gln Leu Arg Leu Val Lys
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Ser Gln Leu Gln Gly Leu Ser Ala Ser Ala Gly Leu Leu Glu Gln Met
225                 230                 235                 240
Arg His Met Glu Thr Gln Ala Lys Asp Leu Arg Asn Gln Leu Leu Asn
                245                 250                 255
Tyr Arg Ser Ala Ile Ser Asn His Gly Ser Lys Ile Glu Gly Leu Glu
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Arg Glu Leu Thr Asp Leu Asn Gln Glu Phe Glu Thr Leu Gln Glu Lys
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Arg Ala Thr Gln Ser Ala Lys Glu Leu Asp Val Lys Ile Lys Asn Val
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Ile Arg Asn Val His Ile Leu Leu Lys Gln Ile Ser Gly Thr Asp Gly
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Glu Gly Asn Asn Val Pro Ser Gly Asp Phe Ser Arg Glu Trp Ala Glu
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Ala Gln Arg Met Met Arg Glu Leu Arg Asn Arg Asn Phe Gly Lys His
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    370                 375                 380
Arg Ile Arg Thr Trp Gln Lys Thr His Gln Gly Glu Asn Asn Gly Leu
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Asp Leu Arg Ala Arg Leu Gln Glu Ala Ala Ala Gln Ala Lys Gln Ala
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Gln Val Lys Glu Ile Asn Ser Leu Gln Ser Asp Phe Thr Lys Tyr Leu
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Thr Thr Ala Asp Ser Ser Leu Leu Gln Thr Asn Ile Ala Leu Gln Leu
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Met Glu Lys Ser Gln Lys Glu Tyr Glu Lys Leu Ala Ala Ser Leu Asn
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Glu Ala Arg Gln Glu Leu Ser Asp Lys Val Arg Glu Leu Ser Arg Ser
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Ala Gly Lys Thr Ser Leu Val Glu Glu Ala Glu Lys His Ala Arg Ser
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Gly Asp Glu Leu Val Arg Cys Ala Val Asp Ala Ala Thr Ala Tyr Glu
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Asn Ile Leu Asn Ala Ile Lys Ala Ala Glu Asp Ala Ala Asn Arg Ala
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Ala Ser Ala Ser Glu Ser Ala Leu Gln Thr Val Ile Lys Glu Asp Leu
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Pro Arg Lys Ala Lys Thr Leu Ser Ser Asn Ser Asp Lys Leu Leu Asn
        595                 600                 605
Glu Ala Lys Met Thr Gln Lys Lys Leu Lys Gln Glu Val Ser Pro Ala
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Leu Asn Asn Leu Gln Gln Thr Leu Asn Ile Val Thr Val Gln Lys Glu
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Val Ile Asp Thr Asn Leu Thr Thr Leu Arg Asp Gly Leu His Gly Ile
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Gln Arg Gly Asp Ile Asp Ala Met Ile Ser Ser Ala Lys Ser Met Val
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Arg Lys Ala Asn Asp Ile Thr Asp Glu Val Leu Asp Gly Leu Asn Pro
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Ile Gln Thr Asp Val Glu Arg Ile Lys Asp Thr Tyr Gly Arg Thr Gln
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Asn Glu Asp Phe Lys Lys Ala Leu Thr Asp Ala Asp Asn Ser Val Asn
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Lys Leu Thr Asn Lys Leu Pro Asp Leu Trp Arg Lys Ile Glu Ser Ile
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Asn Gln Gln Leu Leu Pro Leu Gly Asn Ile Ser Asp Asn Met Asp Arg
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Gln Arg Pro Asn Ser Arg Glu Asn Gly Gly Thr Glu Asn Met Phe Val
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Met Tyr Leu Gly Asn Lys Asp Ala Ser Arg Asp Tyr Ile Gly Met Ala
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Val Val Asp Gly Gln Leu Thr Cys Val Tyr Asn Leu Gly Asp Arg Glu
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Ala Glu Leu Gln Val Asp Gln Ile Leu Thr Lys Ser Glu Thr Lys Glu
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Ala Val Met Asp Arg Val Lys Phe Gln Arg Ile Tyr Gln Phe Ala Arg
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Leu Asn Tyr Thr Lys Gly Ala Thr Ser Ser Lys Pro Glu Thr Pro Gly
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Val Tyr Asp Met Asp Gly Arg Asn Ser Asn Thr Leu Leu Asn Leu Asp
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Pro Glu Asn Val Val Phe Tyr Val Gly Gly Tyr Pro Pro Asp Phe Lys
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Leu Pro Ser Arg Leu Ser Phe Pro Pro Tyr Lys Gly Cys Ile Glu Leu
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Asp Asp Leu Asn Glu Asn Val Leu Ser Leu Tyr Asn Phe Lys Lys Thr
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Phe Asn Leu Asn Thr Thr Glu Val Glu Pro Cys Arg Arg Arg Lys Glu
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Glu Ser Asp Lys Asn Tyr Phe Glu Gly Thr Gly Tyr Ala Arg Val Pro
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Thr Gln Pro His Ala Pro Ile Pro Thr Phe Gly Gln Thr Ile Gln Thr
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Phe Asp Phe Ser Thr Tyr Tyr Leu Gly Gly Ile Pro Ile Ala Ile Arg
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Ile Leu Leu Asp His Gln Thr Trp Thr Arg Asn Leu Gln Val Thr Leu
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Phe Lys Ser Pro Gln Thr Tyr Met Asp Gly Leu Leu His Tyr Val Ser
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aaagcagagc acaagaagca ctgagtatgg gcaatgccac tttttatgaa gttgagagca 2940
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aagccatgaa gagactctcc tacatcagcc agaaggtttc agatgccagt gacaagaccc 3060
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ccggggaggc cctggaaatc tccagtgaga ttgaacagga gattgggagt ctgaacttgg 3180
aagccaatgt gacagcagat ggagccttgg ccatggaaaa gggactggcc tctctgaaga 3240
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agcctctcag tgtagatgaa gaggggctgg tcttactgga gcagaagctt cccgagcca 3480
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agaacttgga gaacattagg gacaacctgc ccccaggctg ctacaatacc caggctcttg 3660
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atgatcaagg atctggaccc caaagaatag actggatgga aagacaaact gcacaggcag 3960
atgtttgcct cataatagtc gtaagtggag tcctggaatt tggacaagtg ctgttgggat 4020
atagtcaact tattctttga gtaatgtgac taaaggaaaa aactttgact ttgcccaggc 4080
atgaaattct tcctaatgtc agaacagagt gcaacccagt cacactgtgg ccagtaaaat 4140
actattgcct catattgtcc tctgcaagct tcttgctgat cagagttcct cctacttaca 4200
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attagtccta attcaatcct acttttcgaa caccaaaaat gatgcgcatc aatgtatttt 4560
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gcaataaccg cttggtttgc aacctctttg ctcaacagaa catatgttgc aagaccctcc 4920
catgggggca cttgagtttt ggcaaggctg acagagctct gggttgtgca catttctttg 4980
cattccagct gtcactctgt gcctttctac aactgattgc aacagactgt tgagttatga 5040
taacaccagt gggaattgct ggaggaacca gaggcacttc caccttggct gggaagacta 5100
tggtgctgcc ttgcttctgt atttccttgg attttcctga aagtgttttt aaataaagaa 5160
caattgttag atgcc                                                  5175
```

<210> 115
<211> 1193
<212> PRT
<213> Homo sapiens

<400> 115

```
Met Pro Ala Leu Trp Leu Gly Cys Cys Leu Cys Phe Ser Leu Leu Leu
1               5                   10                  15
Pro Ala Ala Arg Ala Thr Ser Arg Arg Glu Val Cys Asp Cys Asn Gly
                20                  25                  30
Lys Ser Arg Gln Cys Ile Phe Asp Arg Glu Leu His Arg Gln Thr Gly
            35                  40                  45
Asn Gly Phe Arg Cys Leu Asn Cys Asn Asp Asn Thr Asp Gly Ile His
        50                  55                  60
Cys Glu Lys Cys Lys Asn Gly Phe Tyr Arg His Arg Glu Arg Asp Arg
65                  70                  75                  80
Cys Leu Pro Cys Asn Cys Asn Ser Lys Gly Ser Leu Ser Ala Arg Cys
                85                  90                  95
```

```
Asp Asn Ser Gly Arg Cys Ser Cys Lys Pro Gly Val Thr Gly Ala Arg
            100             105             110
Cys Asp Arg Cys Leu Pro Gly Phe His Met Leu Thr Asp Ala Gly Cys
        115             120             125
Thr Gln Asp Gln Arg Leu Leu Asp Ser Lys Cys Asp Cys Asp Pro Ala
    130             135             140
Gly Ile Ala Gly Pro Cys Asp Ala Gly Arg Cys Val Cys Lys Pro Ala
145             150             155             160
Val Thr Gly Glu Arg Cys Asp Arg Cys Arg Ser Gly Tyr Tyr Asn Leu
            165             170             175
Asp Gly Gly Asn Pro Glu Gly Cys Thr Gln Cys Phe Cys Tyr Gly His
            180             185             190
Ser Ala Ser Cys Arg Ser Ser Ala Glu Tyr Ser Val His Lys Ile Thr
        195             200             205
Ser Thr Phe His Gln Asp Val Asp Gly Trp Lys Ala Val Gln Arg Asn
    210             215             220
Gly Ser Pro Ala Lys Leu Gln Trp Ser Gln Arg His Gln Asp Val Phe
225             230             235             240
Ser Ser Ala Gln Arg Leu Asp Pro Val Tyr Phe Val Ala Pro Ala Lys
            245             250             255
Phe Leu Gly Asn Gln Gln Val Ser Tyr Gly Gln Ser Leu Ser Phe Asp
            260             265             270
Tyr Arg Val Asp Arg Gly Gly Arg His Pro Ser Ala His Asp Val Ile
        275             280             285
Leu Glu Gly Ala Gly Leu Arg Ile Thr Ala Pro Leu Met Pro Leu Gly
        290             295             300
Lys Thr Leu Pro Cys Gly Leu Thr Lys Thr Tyr Thr Phe Arg Leu Asn
305             310             315             320
Glu His Pro Ser Asn Asn Trp Ser Pro Gln Leu Ser Tyr Phe Glu Tyr
            325             330             335
Arg Arg Leu Leu Arg Asn Leu Thr Ala Leu Arg Ile Arg Ala Thr Tyr
        340             345             350
Gly Glu Tyr Ser Thr Gly Tyr Ile Asp Asn Val Thr Leu Ile Ser Ala
        355             360             365
Arg Pro Val Ser Gly Ala Pro Ala Pro Trp Val Glu Gln Cys Ile Cys
    370             375             380
Pro Val Gly Tyr Lys Gly Gln Phe Cys Gln Asp Cys Ala Ser Gly Tyr
385             390             395             400
Lys Arg Asp Ser Ala Arg Leu Gly Pro Phe Gly Thr Cys Ile Pro Cys
            405             410             415
Asn Cys Gln Gly Gly Gly Ala Cys Asp Pro Asp Thr Gly Asp Cys Tyr
            420             425             430
Ser Gly Asp Glu Asn Pro Asp Ile Glu Cys Ala Asp Cys Pro Ile Gly
        435             440             445
Phe Tyr Asn Asp Pro His Asp Pro Arg Ser Cys Lys Pro Cys Pro Cys
    450             455             460
His Asn Gly Phe Ser Cys Ser Val Met Pro Glu Thr Glu Glu Val Val
465             470             475             480
Cys Asn Asn Cys Pro Pro Gly Val Thr Gly Ala Arg Cys Glu Leu Cys
            485             490             495
Ala Asp Gly Tyr Phe Gly Asp Pro Phe Gly Glu His Gly Pro Val Arg
        500             505             510
Pro Cys Gln Pro Cys Gln Cys Asn Asn Asn Val Asp Pro Ser Ala Ser
        515             520             525
Gly Asn Cys Asp Arg Leu Thr Gly Arg Cys Leu Lys Cys Ile His Asn
530             535             540
Thr Ala Gly Ile Tyr Cys Asp Gln Cys Lys Ala Gly Tyr Phe Gly Asp
545             550             555             560
Pro Leu Ala Pro Asn Pro Ala Asp Lys Cys Arg Ala Cys Asn Cys Asn
```

```
                     565                     570                     575
Pro Met Gly Ser Glu Pro Val Gly Cys Arg Ser Asp Gly Thr Cys Val
            580                     585                     590
Cys Lys Pro Gly Phe Gly Gly Pro Asn Cys Glu His Gly Ala Phe Ser
            595                     600                     605
Cys Pro Ala Cys Tyr Asn Gln Val Lys Ile Gln Met Asp Gln Phe Met
        610                     615                     620
Gln Gln Leu Gln Arg Met Glu Ala Leu Ile Ser Lys Ala Gln Gly Gly
625                     630                     635                     640
Asp Gly Val Val Pro Asp Thr Glu Leu Glu Gly Arg Met Gln Gln Ala
                    645                     650                     655
Glu Gln Ala Leu Gln Asp Ile Leu Arg Asp Ala Gln Ile Ser Glu Gly
            660                     665                     670
Ala Ser Arg Ser Leu Gly Leu Gln Leu Ala Lys Val Arg Ser Gln Glu
            675                     680                     685
Asn Ser Tyr Gln Ser Arg Leu Asp Asp Leu Lys Met Thr Val Glu Arg
        690                     695                     700
Val Arg Ala Leu Gly Ser Gln Tyr Gln Asn Arg Val Arg Asp Thr His
705                     710                     715                     720
Arg Leu Ile Thr Gln Met Gln Leu Ser Leu Ala Glu Ser Glu Ala Ser
            725                     730                     735
Leu Gly Asn Thr Asn Ile Pro Ala Ser Asp His Tyr Val Gly Pro Asn
            740                     745                     750
Gly Phe Lys Ser Leu Ala Gln Glu Ala Thr Arg Leu Ala Glu Ser His
            755                     760                     765
Val Glu Ser Ala Ser Asn Met Glu Gln Leu Thr Arg Glu Thr Glu Asp
        770                     775                     780
Tyr Ser Lys Gln Ala Leu Ser Leu Val Arg Lys Ala Leu His Glu Gly
785                     790                     795                     800
Val Gly Ser Gly Ser Gly Ser Pro Asp Gly Ala Val Val Gln Gly Leu
            805                     810                     815
Val Glu Lys Leu Glu Lys Thr Lys Ser Leu Ala Gln Gln Leu Thr Arg
            820                     825                     830
Glu Ala Thr Gln Ala Glu Ile Glu Ala Asp Arg Ser Tyr Gln His Ser
            835                     840                     845
Leu Arg Leu Leu Asp Ser Val Ser Arg Leu Gln Gly Val Ser Asp Gln
        850                     855                     860
Ser Phe Gln Val Glu Glu Ala Lys Arg Ile Lys Gln Lys Ala Asp Ser
865                     870                     875                     880
Leu Ser Thr Leu Val Thr Arg His Met Asp Glu Phe Lys Arg Thr Gln
            885                     890                     895
Lys Asn Leu Gly Asn Trp Lys Glu Glu Ala Gln Gln Leu Leu Gln Asn
            900                     905                     910
Gly Lys Ser Gly Arg Glu Lys Ser Asp Gln Leu Leu Ser Arg Ala Asn
            915                     920                     925
Leu Ala Lys Ser Arg Ala Gln Glu Ala Leu Ser Met Gly Asn Ala Thr
        930                     935                     940
Phe Tyr Glu Val Glu Ser Ile Leu Lys Asn Leu Arg Glu Phe Asp Leu
945                     950                     955                     960
Gln Val Asp Asn Arg Lys Ala Glu Ala Glu Ala Met Lys Arg Leu
            965                     970                     975
Ser Tyr Ile Ser Gln Lys Val Ser Asp Ala Ser Asp Lys Thr Gln Gln
        980                     985                     990
Ala Glu Arg Ala Leu Gly Ser Ala Ala Ala Asp Ala Gln Arg Ala Lys
        995                     1000                    1005
Asn Gly Ala Gly Glu Ala Leu Glu Ile Ser Ser Glu Ile Glu Gln Glu
    1010                    1015                    1020
Ile Gly Ser Leu Asn Leu Glu Ala Asn Val Thr Ala Asp Gly Ala Leu
1025                    1030                    1035                    1040
```

```
Ala Met Glu Lys Gly Leu Ala Ser Leu Lys Ser Glu Met Arg Glu Val
              1045                1050                1055
Glu Gly Glu Leu Glu Arg Lys Glu Leu Glu Phe Asp Thr Asn Met Asp
              1060                1065                1070
Ala Val Gln Met Val Ile Thr Glu Ala Gln Lys Val Asp Thr Arg Ala
         1075                1080                1085
Lys Asn Ala Gly Val Thr Ile Gln Asp Thr Leu Asn Thr Leu Asp Gly
         1090                1095                1100
Leu Leu His Leu Met Asp Gln Pro Leu Ser Val Asp Glu Glu Gly Leu
1105                1110                1115                1120
Val Leu Leu Glu Gln Lys Leu Ser Arg Ala Lys Thr Gln Ile Asn Ser
              1125                1130                1135
Gln Leu Arg Pro Met Met Ser Glu Leu Glu Glu Arg Ala Arg Gln Gln
              1140                1145                1150
Arg Gly His Leu His Leu Leu Glu Thr Ser Ile Asp Gly Ile Leu Ala
         1155                1160                1165
Asp Val Lys Asn Leu Glu Asn Ile Arg Asp Asn Leu Pro Pro Gly Cys
         1170                1175                1180
Tyr Asn Thr Gln Ala Leu Glu Gln Gln
1185                1190
```

```
<210> 116
<211> 749
<212> DNA
<213> Homo sapiens

<400> 116
atggcggcta acgctactac caacccgtcg cagctgctgc cgttagagct tgtggacaaa 60
tgtatacggat caagaattca catcgtgatg aagagtgata aggaaattgt tggtactctt 120
ctaggatttg atgactttgt caatatggta ctggaagatg tcactgagtt tgaaatcaca 180
ccagaaggaa gaaggattac taaattagat cagattttgc taaatggaaa taatataaca 240
atgctggttc ctggaggaga aggacctgaa gtgtgaatga gtttccttga cttacactag 300
attttgtttt ggcttataat gacaagaaaa tggaattttt tttcccactt tctaatgttt 360
aaatcccata aagctaagtt tcccgttaaa gggaagtgct ttgaagatgt gtacccattt 420
ttgtaagtta atcatgatta tcctggaaaa agaagaaaag aacttcttct tttgcagatg 480
aaaataaagg tgttttttggt taactgtcat tttgtttatt ctactgcagt agccagtgga 540
acaaagtttg tagttatttt gccacttact tttctgtcat tatatgctta tttgtttttgt 600
catttacgtg accatttgat tctcaaacaa aagttgttcc aaacaaatg atgaactttg 660
atttgaacag gtgcatttaa acaaccggaa atgatcactt agaaaattca attaaaatgc 720
tgttgttttg taaaaaaaaa aaaaaaaaa                                    749
```

```
<210> 117
<211> 91
<212> PRT
<213> Homo sapiens

<400> 117
Met Ala Ala Asn Ala Thr Thr Asn Pro Ser Gln Leu Leu Pro Leu Glu
1                   5                   10                  15
Leu Val Asp Lys Cys Ile Gly Ser Arg Ile His Ile Val Met Lys Ser
              20                  25                  30
Asp Lys Glu Ile Val Gly Thr Leu Leu Gly Phe Asp Asp Phe Val Asn
         35                  40                  45
Met Val Leu Glu Asp Val Thr Glu Phe Glu Ile Thr Pro Glu Gly Arg
    50                  55                  60
Arg Ile Thr Lys Leu Asp Gln Ile Leu Leu Asn Gly Asn Asn Ile Thr
65                  70                  75                  80
Met Leu Val Pro Gly Gly Glu Gly Pro Glu Val
```

85   90

<210> 118
<211> 1717
<212> DNA
<213> Homo sapiens

<400> 118
gtggattctt gtccatagtg catctgcttt aagaattaac gaaagcagtg tcaagacagt 60
aaggattcaa accatttgcc aaaaatgagt ctaagtgcat ttactctctt cctggcattg 120
attggtggta ccagtggcca gtactatgat tatgattttc ccctatcaat ttatgggcaa 180
tcatcaccaa actgtgcacc agaatgtaac tgccctgaaa gctacccaag tgccatgtac 240
tgtgatgagc tgaaattgaa aagtgtacca atggtgcctc ctggaatcaa gtatctttac 300
cttaggaata ccagattga ccatattgat gaaaaggcct ttgagaatgt aactgatctg 360
cagtggctca ttctagatca caaccttcta gaaaactcca agataaaagg gagagttttc 420
tctaaattga aacaactgaa gaagctgcat ataaccaca caaccctgac agagtctgtg 480
ggcccacttc ccaaatctct ggaggatctg cagcttactc ataacaagat cacaaagctg 540
ggctcttttg aaggattggt aaacctgacc ttcatccatc tccagcacaa tcggctgaaa 600
gaggatgctg tttcagctgc ttttaaaggt cttaaatcac tcgaatacct tgacttgagc 660
ttcaatcaga tagccagact gccttctggt ctccctgtct ctcttctaac tctctactta 720
gacaacaata agatcagcaa catccctgat gagtatttca agcgttttaa tgcattgcag 780
tatctgcgtt tatctcacaa cgaactggct gatagtggaa tacctggaaa ttctttcaat 840
gtgtcatccc tggttgagct ggatctgtcc tataacaagc ttaaaaacat accaactgtc 900
aatgaaaacc ttgaaaacta ttacctggag gtcaatcaac ttgagaagtt tgacataaag 960
agcttctgca agatcctggg gccattatcc tactccaaga tcaagcattt gcgtttggat 1020
ggcaatcgca tctcagaaac cagtcttcca ccggatatgt atgaatgtct acgtgttgct 1080
aacgaagtca ctcttaatta atatctgtat cctggaacaa tattttatgg ttatgttttt 1140
ctgtgtgtca gtttttcatag tatccatatt ttattactgt ttattacttc catgaatttt 1200
aaaatctgag ggaaatgttt tgtaaacatt tattttttttt aaagaaaaga tgaaaggcag 1260
gcctatttca tcacaagaac acacacatat acacgaatag acatcaaact caatgcttta 1320
tttgtaaatt tagtgtttttt ttatttctac tgtcaaatga tgtgcaaaac cttttttactgg 1380
ttgcatggaa atcagccaag ttttataatc cttaaatctt aatgttcctc aaagcttgga 1440
ttaaatacat atggatgtta ctctcttgca ccaaattatc ttgatacatt caaatttgtc 1500
tggtaaaaaa ataggtggta gatattgagg ccaagaatat tgcaaaatac atgaagcttc 1560
atgcacttaa agaagtattt ttagaataag aatttgcata cttacctagt gaaactttc 1620
tagaattatt tttcactcta agtcatgtat gtttctcttt gattatttgc atgttatgtt 1680
taataagcta ctagcaaaat aaaacatagc aaatggc   1717

<210> 119
<211> 338
<212> PRT
<213> Homo sapiens

<400> 119
Met Ser Leu Ser Ala Phe Thr Leu Phe Leu Ala Leu Ile Gly Gly Thr
  1   5   10   15
Ser Gly Gln Tyr Tyr Asp Tyr Asp Phe Pro Leu Ser Ile Tyr Gly Gln
             20   25   30
Ser Ser Pro Asn Cys Ala Pro Glu Cys Asn Cys Pro Glu Ser Tyr Pro
         35   40   45
Ser Ala Met Tyr Cys Asp Glu Leu Lys Leu Lys Ser Val Pro Met Val
     50   55   60
Pro Pro Gly Ile Lys Tyr Leu Tyr Leu Arg Asn Asn Gln Ile Asp His
 65   70   75   80
Ile Asp Glu Lys Ala Phe Glu Asn Val Thr Asp Leu Gln Trp Leu Ile
                 85   90   95
Leu Asp His Asn Leu Leu Glu Asn Ser Lys Ile Lys Gly Arg Val Phe
            100   105   110

187

```
Ser Lys Leu Lys Gln Leu Lys Lys Leu His Ile Asn His Asn Asn Leu
        115             120                 125
Thr Glu Ser Val Gly Pro Leu Pro Lys Ser Leu Glu Asp Leu Gln Leu
        130             135                 140
Thr His Asn Lys Ile Thr Lys Leu Gly Ser Phe Glu Gly Leu Val Asn
145             150                 155                 160
Leu Thr Phe Ile His Leu Gln His Asn Arg Leu Lys Glu Asp Ala Val
                165                 170                 175
Ser Ala Ala Phe Lys Gly Leu Lys Ser Leu Glu Tyr Leu Asp Leu Ser
            180                 185                 190
Phe Asn Gln Ile Ala Arg Leu Pro Ser Gly Leu Pro Val Ser Leu Leu
            195                 200                 205
Thr Leu Tyr Leu Asp Asn Asn Lys Ile Ser Asn Ile Pro Asp Glu Tyr
        210                 215                 220
Phe Lys Arg Phe Asn Ala Leu Gln Tyr Leu Arg Leu Ser His Asn Glu
225                 230                 235                 240
Leu Ala Asp Ser Gly Ile Pro Gly Asn Ser Phe Asn Val Ser Ser Leu
            245                 250                 255
Val Glu Leu Asp Leu Ser Tyr Asn Lys Leu Lys Asn Ile Pro Thr Val
            260                 265                 270
Asn Glu Asn Leu Glu Asn Tyr Tyr Leu Glu Val Asn Gln Leu Glu Lys
        275                 280                 285
Phe Asp Ile Lys Ser Phe Cys Lys Ile Leu Gly Pro Leu Ser Tyr Ser
    290                 295                 300
Lys Ile Lys His Leu Arg Leu Asp Gly Asn Arg Ile Ser Glu Thr Ser
305                 310                 315                 320
Leu Pro Pro Asp Met Tyr Glu Cys Leu Arg Val Ala Asn Glu Val Thr
                325                 330                 335
Leu Asn
```

```
<210> 120
<211> 1334
<212> DNA
<213> Homo sapiens

<400> 120
gcagaccccc atcatgggca gccagagctc caaggctccc cggggcgacg tgaccgccga 60
ggaggcagca ggcgcttccc ccgcgaaggc caacggccag gagaatggcc acgtgaaaag 120
caatggagac ttatccccca agggtgaagg ggagtcgccc cctgtgaacg gaacagatga 180
ggcagccggg gccactggcg atgccatcga gccagcaccc cctagccagg gtgctgaggc 240
caagggggag gtcccccca aggagacccc caagaagaag aagaaattct ctttcaagaa 300
gcctttcaaa ttgagcggcc tgtccttcaa gagaaatcgg aaggagggtg ggggtgattc 360
ttctgcctcc tcacccacag aggaagagca ggagcagggg gagatcggtg cctgcagcga 420
cgagggcact gctcaggaag ggaaggccgc agccacccct gagagccagg aaccccaggc 480
caagggggca gaggctagtg cagcctcaga agaagaggca gggccccagg ctacagagcc 540
atccactccc tcggggccgg agagtggccc tacaccagcc agcgctgagc agaatgagta 600
gctaggtagg ggcaggtggg tgatctctaa gctgcaaaaa ctgtgctgtc cttgtgaggt 660
cactgcctgg acctggtgcc ctggctgcct tcctgtgccc agaaaggaag gggctattgc 720
ctcctcccag ccacgttccc tttcctcctc tccctcctgt ggattctccc atcagccatc 780
tggttctcct cttaaggcca gttgaagatg gtcccttaca gcttcccaag ttaggttagt 840
gatgtgaaat gctcctgtcc ctggccctac ctccttccct gtccccaccc ctgcataagg 900
cagttgttgg ttttcttccc caattctttt ccaagtaggt tttgtttacc ctactcccca 960
aatccctgag ccagaagtgg ggtgcttata ctcccaaacc ttgagtgtcc agccttcccc 1020
tgttgttttt agtctcttgt gctgtgccta gtggcacctg ggctggggag gacactgccc 1080
cgtctaggtt tttataaatg tcttactcaa gttcaaacct ccagcctgtg aatcaactgt 1140
gtctcttttt tgacttggta agcaagtatt aggctttggg gtggggggag gtctgtaatg 1200
tgaaacaact tcttgtcttt ttttctccca ctgttgtaaa taactttaa tggccaaacc 1260
```

ccagatttgt actttttttt tttttctaac tgctaaaacc attctcttcc acctggtttt 1320
actgtaacat ttgg   1334

<210> 121
<211> 195
<212> PRT
<213> Homo sapiens

<400> 121
Met Gly Ser Gln Ser Ser Lys Ala Pro Arg Gly Asp Val Thr Ala Glu
1   5   10   15
Glu Ala Ala Gly Ala Ser Pro Ala Lys Ala Asn Gly Gln Glu Asn Gly
            20   25   30
His Val Lys Ser Asn Gly Asp Leu Ser Pro Lys Gly Glu Gly Glu Ser
        35   40   45
Pro Pro Val Asn Gly Thr Asp Glu Ala Ala Gly Ala Thr Gly Asp Ala
    50   55   60
Ile Glu Pro Ala Pro Pro Ser Gln Gly Ala Glu Ala Lys Gly Glu Val
65   70   75   80
Pro Pro Lys Glu Thr Pro Lys Lys Lys Lys Phe Ser Phe Lys Lys
                85   90   95
Pro Phe Lys Leu Ser Gly Leu Ser Phe Lys Arg Asn Arg Lys Glu Gly
            100   105   110
Gly Gly Asp Ser Ser Ala Ser Ser Pro Thr Glu Glu Glu Gln Glu Gln
        115   120   125
Gly Glu Ile Gly Ala Cys Ser Asp Glu Gly Thr Ala Gln Glu Gly Lys
    130   135   140
Ala Ala Ala Thr Pro Glu Ser Gln Glu Pro Gln Ala Lys Gly Ala Glu
145   150   155   160
Ala Ser Ala Ala Ser Glu Glu Glu Ala Gly Pro Gln Ala Thr Glu Pro
            165   170   175
Ser Thr Pro Ser Gly Pro Glu Ser Gly Pro Thr Pro Ala Ser Ala Glu
        180   185   190
Gln Asn Glu
    195

<210> 122
<211> 1081
<212> DNA
<213> Homo sapiens

<400> 122
attgcaactt ggtctcacag tggcttaggc cagggtggga gcagtgaacg gagtcacaaa 60
agaaattttt cagctgtcct ctctgacacc accccggcct gcctctttgt tgccatgaga 120
gctgcctacc tcttcctgct attcctgcct gcaggcttgc tggctcaggg ccagtatgat 180
ctggacccgc tgccgccgtt ccctgaccac gtccagtaca cccactatag cgaccagatc 240
gacaacccag actactatga ttatcaagag gtgactcctc ggccctccga ggaacagttc 300
cagttccagt cccagcagca agtccaacag gaagtcatcc cagccccaac cccagaacca 360
ggaaatgcag agctggagcc cacagagcct gggcctcttg actgccgtga ggaacagtac 420
ccgtgcaccc gcctctactc catacacagg ccttgcaaac agtgtctcaa cgaggtctgc 480
ttctacagcc tccgccgtgt gtacgtcatt aacaaggaga tctgtgttcg tacagtgtgt 540
gcccacgagg agctcctccg agctgacctc tgtcgggaca agttctccaa atgtggcgtg 600
atggccagca gcggcctgtg ccaatccgtg gcggcctcct gtgccaggag ctgtgggagc 660
tgctagggtg gtgctggcat cctgagtcct ggcctcctg ggatctgggg ccctcgggct 720
acctgacctg gtgcttttt ccccatcccc atgttccttt tattctgaaa aagttagtgg 780
actgcagccc tggggggttgc aggctgcggt gcctcaggcc cctccttcag cctgtggcca 840
cctctggggc acgatggggg ctccccactg cccagtctgc ccctcgggtt gggggagtat 900
cccaggcctc tctgtgggac ctgggcctga cgggcccttc tcagcccgtt ttgaggacag 960

acagtccccc gaggtaggct acatcccccc accccagctg gtctgcttgg atttcctaca 1020
gcccccgtgg gcatggacca cctttatttt atacaaaatt aaaaacaagt ttttacaaaa 1080
a   1081

<210> 123
<211> 183
<212> PRT
<213> Homo sapiens

<400> 123
Met Arg Ala Ala Tyr Leu Phe Leu Leu Phe Leu Pro Ala Gly Leu Leu
1   5   10   15
Ala Gln Gly Gln Tyr Asp Leu Asp Pro Leu Pro Pro Phe Pro Asp His
            20   25   30
Val Gln Tyr Thr His Tyr Ser Asp Gln Ile Asp Asn Pro Asp Tyr Tyr
        35   40   45
Asp Tyr Gln Glu Val Thr Pro Arg Pro Ser Glu Glu Gln Phe Gln Phe
    50   55   60
Gln Ser Gln Gln Gln Val Gln Gln Glu Val Ile Pro Ala Pro Thr Pro
65   70   75   80
Glu Pro Gly Asn Ala Glu Leu Glu Pro Thr Glu Pro Gly Pro Leu Asp
                85   90   95
Cys Arg Glu Glu Gln Tyr Pro Cys Thr Arg Leu Tyr Ser Ile His Arg
            100   105   110
Pro Cys Lys Gln Cys Leu Asn Glu Val Cys Phe Tyr Ser Leu Arg Arg
        115   120   125
Val Tyr Val Ile Asn Lys Glu Ile Cys Val Arg Thr Val Cys Ala His
    130   135   140
Glu Glu Leu Leu Arg Ala Asp Leu Cys Arg Asp Lys Phe Ser Lys Cys
145   150   155   160
Gly Val Met Ala Ser Ser Gly Leu Cys Gln Ser Val Ala Ala Ser Cys
                165   170   175
Ala Arg Ser Cys Gly Ser Cys
            180

<210> 124
<211> 1066
<212> DNA
<213> Homo sapiens

<400> 124
ggccaagggg cggctccggc gggcggactc ggagcgggcg gcggagtgac ccggacagct 60
gtcctctctg acaccacccc ggcctgcctc tttgttgcca tgagagctgc ctacctcttc 120
ctgctattcc tgcctgcagg cttgctggct cagggccagt atgatctgga cccgctgccg 180
ccgttccctg accacgtcca gtacacccac tatagcgacc agatcgacaa cccagactac 240
tatgattatc aagaggtgac tcctcggccc tccgaggaac agttccagtt ccagtcccag 300
cagcaagtcc aacaggaagt catcccagcc ccaaccccag aaccaggaaa tgcagagctg 360
gagcccacag agcctgggcc tcttgactgc cgtgaggaac agtacccgtg cacccgcctc 420
tactccatac acaggccttg caaacagtgt ctcaacgagg tctgcttcta cagcctccgc 480
cgtgtgtacg tcattaacaa ggagatctgt gttcgtacag tgtgtgccca cgaggagctc 540
ctccgagctg acctctgtcg ggacaagttc tccaaatgtg gcgtgatggc cagcagcggc 600
ctgtgccaat ccgtggcggc ctcctgtgcc aggagctgtg ggagctgcta gggtggtgct 660
ggcatcctga gtcctggccc tcctgggatc tggggccctc gggctacctg acctggtgct 720
tttttcccca tccccatgtt cctttattc tgaaaaagtt agtggactgc agccctgggg 780
gttgcaggct gcggtgcctc aggcccctcc ttcagcctgt ggccacctct ggggcacgat 840
gggggctccc cactgcccag tctgcccctc gggttggggg agtatcccag gcctctctgt 900
gggacctggg cctgacgggc ccttctcagc ccgttttgag gacagacagt ccccgaggt 960
aggctacatc ccccaccccc agctggtctg cttggatttc ctacagcccc cgtgggcatg 1020

gaccaccttt attttataca aaattaaaaa caagttttta caaaaa　　　　　1066

<210> 125
<211> 183
<212> PRT
<213> Homo sapiens

<400> 125

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Met Arg Ala Ala Tyr Leu Phe Leu Leu Phe Leu Pro Ala Gly Leu Leu
1               5                   10                  15
Ala Gln Gly Gln Tyr Asp Leu Asp Pro Leu Pro Pro Phe Pro Asp His
            20                  25                  30
Val Gln Tyr Thr His Tyr Ser Asp Gln Ile Asp Asn Pro Asp Tyr Tyr
        35                  40                  45
Asp Tyr Gln Glu Val Thr Pro Arg Pro Ser Glu Glu Gln Phe Gln Phe
    50                  55                  60
Gln Ser Gln Gln Gln Val Gln Gln Glu Val Ile Pro Ala Pro Thr Pro
65                  70                  75                  80
Glu Pro Gly Asn Ala Glu Leu Glu Pro Thr Glu Pro Gly Pro Leu Asp
                85                  90                  95
Cys Arg Glu Glu Gln Tyr Pro Cys Thr Arg Leu Tyr Ser Ile His Arg
            100                 105                 110
Pro Cys Lys Gln Cys Leu Asn Glu Val Cys Phe Tyr Ser Leu Arg Arg
        115                 120                 125
Val Tyr Val Ile Asn Lys Glu Ile Cys Val Arg Thr Val Cys Ala His
    130                 135                 140
Glu Glu Leu Leu Arg Ala Asp Leu Cys Arg Asp Lys Phe Ser Lys Cys
145                 150                 155                 160
Gly Val Met Ala Ser Ser Gly Leu Cys Gln Ser Val Ala Ala Ser Cys
                165                 170                 175
Ala Arg Ser Cys Gly Ser Cys
                180
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<210> 126
<211> 1611
<212> DNA
<213> Homo sapiens

<400> 126

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gatcatggtc tggatctgca acacgggcca ggccaaagtc acagatcttg agatcacagg 120
tggtgttgag cagcaggcag gcaggcaatc ggtccgagtg gctgtcggct cttcagctct 180
ccgctcggcg tcttccttcc tctcccggtc agcgtcggcg gctgcaccgg cggcgggcag 240
tcctgcggga ggggcgacaa gagctgaggc gcggccgccg agcgtcgagc tcagcgcggc 300
ggaggcggcg gcggcccggc agccaacatg gcggcggcgg cggcggcggg cgcgggcccg 360
gagatggtcc gcgggcaggt gttcgacgtg gggccgcgct acaccaacct ctcgtacatc 420
ggcgagggcg cctacggcat ggtgtgctct gcttatgata atgtcaacaa agttcgagta 480
gctatcaaga aaatcagccc ctttgagcac cagacctact gccagagaac cctgagggag 540
ataaaaatct tactgcgctt cagacatgag aacatcattg gaatcaatga cattattcga 600
gcaccaacca tcgagcaaat gaaagatgta tatatagtac aggacctcat ggaaacagat 660
ctttacaagc tcttgaagac acaacacctc agcaatgacc atatctgcta ttttctctac 720
cagatcctca gagggttaaa atatatccat tcagctaacg ttctgcaccg tgacctcaag 780
ccttccaacc tgctgctcaa caccacctgt gatctcaaga tctgtgactt ggcctggcc 840
cgtgttgcag atccagacca tgatcacaca gggttcctga cagaatatgt ggccacacgt 900
tggtacaggg ctccagaaat tatgttgaat tccaagggct acaccaagtc cattgatatt 960
tggtctgtag ctgcattct ggcagaaatg ctttccaaca ggcccatctt tccagggaag 1020
cattatcttg accagctgaa tcacattttg ggtattcttg atccccatc acaagaagac 1080
ctgaattgta taataaattt aaaagctagg aactatttgc tttctcttcc acacaaaaat 1140
```

```
aaggtgccat ggaacaggct gttcccaaat gctgactcca aagctctgga cttattggac 1200
aaaatgttga cattcaaccc acacaagagg attgaagtag aacaggctct ggcccaccca 1260
tatctggagc agtattacga cccgagtgac gagcccatcg ccgaagcacc attcaagttc 1320
gacatggaat tggatgactt gcctaaggaa aagctaaaag aactaatttt tgaagagact 1380
gctagattcc agccaggata cagatcttaa atttgtcagg acaagggctc agaggactgg 1440
acgtgctcag acatcggtgt tcttcttccc agttcttgac ccctggtcct gtctccagcc 1500
cgtcttggct tatccacttt gactcctttg agccgtttgg aggggcggtt tctggtagtt 1560
gtggctttta tgctttcaaa gaatttcttc agtccagaga attcactggc c        1611
```

<210> 127
<211> 360
<212> PRT
<213> Homo sapiens

<400> 127

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Met Ala Ala Ala Ala Ala Ala Gly Ala Gly Pro Glu Met Val Arg Gly
1               5                   10                  15
Gln Val Phe Asp Val Gly Pro Arg Tyr Thr Asn Leu Ser Tyr Ile Gly
            20                  25                  30
Glu Gly Ala Tyr Gly Met Val Cys Ser Ala Tyr Asp Asn Val Asn Lys
        35                  40                  45
Val Arg Val Ala Ile Lys Lys Ile Ser Pro Phe Glu His Gln Thr Tyr
    50                  55                  60
Cys Gln Arg Thr Leu Arg Glu Ile Lys Ile Leu Leu Arg Phe Arg His
65                  70                  75                  80
Glu Asn Ile Ile Gly Ile Asn Asp Ile Ile Arg Ala Pro Thr Ile Glu
                85                  90                  95
Gln Met Lys Asp Val Tyr Ile Val Gln Asp Leu Met Glu Thr Asp Leu
            100                 105                 110
Tyr Lys Leu Leu Lys Thr Gln His Leu Ser Asn Asp His Ile Cys Tyr
        115                 120                 125
Phe Leu Tyr Gln Ile Leu Arg Gly Leu Lys Tyr Ile His Ser Ala Asn
    130                 135                 140
Val Leu His Arg Asp Leu Lys Pro Ser Asn Leu Leu Leu Asn Thr Thr
145                 150                 155                 160
Cys Asp Leu Lys Ile Cys Asp Phe Gly Leu Ala Arg Val Ala Asp Pro
                165                 170                 175
Asp His Asp His Thr Gly Phe Leu Thr Glu Tyr Val Ala Thr Arg Trp
            180                 185                 190
Tyr Arg Ala Pro Glu Ile Met Leu Asn Ser Lys Gly Tyr Thr Lys Ser
        195                 200                 205
Ile Asp Ile Trp Ser Val Gly Cys Ile Leu Ala Glu Met Leu Ser Asn
    210                 215                 220
Arg Pro Ile Phe Pro Gly Lys His Tyr Leu Asp Gln Leu Asn His Ile
225                 230                 235                 240
Leu Gly Ile Leu Gly Ser Pro Ser Gln Glu Asp Leu Asn Cys Ile Ile
                245                 250                 255
Asn Leu Lys Ala Arg Asn Tyr Leu Leu Ser Leu Pro His Lys Asn Lys
            260                 265                 270
Val Pro Trp Asn Arg Leu Phe Pro Asn Ala Asp Ser Lys Ala Leu Asp
        275                 280                 285
Leu Leu Asp Lys Met Leu Thr Phe Asn Pro His Lys Arg Ile Glu Val
    290                 295                 300
Glu Gln Ala Leu Ala His Pro Tyr Leu Glu Gln Tyr Tyr Asp Pro Ser
305                 310                 315                 320
Asp Glu Pro Ile Ala Glu Ala Pro Phe Lys Phe Asp Met Glu Leu Asp
                325                 330                 335
Asp Leu Pro Lys Glu Lys Leu Lys Glu Leu Ile Phe Glu Glu Thr Ala
            340                 345                 350
```

Arg Phe Gln Pro Gly Tyr Arg Ser
        355   360

<210> 128
<211> 2917
<212> DNA
<213> Homo sapiens

<400> 128
ggaaaaaagc gacttgtggc ggtcgagcgt ggcgcaggcg aatcctcggc actaagcaaa 60
tatggacctc gcggcggcag cggagccggg cgccggcagc cagcacctgg aggtccgcga 120
cgaggtggcc gagaagtgcc agaaactgtt cctggacttc ttggaggagt ttcagagcag 180
cgatggagaa attaaatact tgcaattagc agaggaactg attcgtcctg agagaaacac 240
attggttgtg agttttgtgg acctggaaca atttaaccag caactttcca ccaccattca 300
agaggagttc tatagagttt acccttacct gtgtcgggcc ttgaaaacat tcgtcaaaga 360
ccgtaaagag atccctcttg ccaaggattt ttatgttgca ttccaagacc tgcctaccag 420
acacaagatt cgagagctca cctcatccag aattggtttg ctcactcgca tcagtgggca 480
ggtggtgcgg actcacccag ttcacccaga gcttgtgagc ggaacttttc tgtgcttgga 540
ctgtcagaca gtgatcaggg atgtagaaca gcagttcaaa tacacacagc caaacatctg 600
ccgaaatcca gtttgtgcca acaggaggag attcttactg gatacaaata aatcaagatt 660
tgttgatttt caaaaggttc gtattcaaga acccaagct gagcttcctc gagggagtat 720
cccccgcagt ttagaagtaa ttttaagggc tgaagctgtg gaatcagctc aagctggtga 780
caagtgtgac tttacaggga cactgattgt tgtgcctgac gtctccaagc ttagcacacc 840
aggagcacgt gcagaaacta attcccgtgt cagtggtgtt gatggatatg agacagaagg 900
cattcgagga ctccgggccc ttggtgttag ggacctttct tataggctgg tctttcttgc 960
ctgctgtgtt gcgccaacca acccaaggtt tggggggaaa gagctcagag atgaggaaca 1020
gacagctgag agcattaaga accaaatgac tgtgaaagaa tgggagaaag tgtttgagat 1080
gagtcaagat aaaaatctat accacaatct ttgtaccagc ctgttcccta ctatacatgg 1140
caatgatgaa gtaaaacggg gtgtcctgct gatgctcttt ggtggcgttc caaagacaac 1200
aggagaaggg acctctcttc gaggggacat aaatgtttgc attgttggtg acccaagtac 1260
agctaagagc caatttctca agcacgtgga ggagttcagc cccagagctg tctacaccag 1320
tggtaaagcg tccagtgctg ctggcttaac agcagctgtt gtgagagatg aagaatctca 1380
tgagtttgtc attgaggctg gagctttgat gttggctgat aatggtgtgt gttgtattga 1440
tgaatttgat aagatgacg tgcgggatca agttgctatt catgaagcta tggaacagca 1500
gaccatatcc atcactaaag caggagtgaa ggctactctg aacgcccgga cgtccatttt 1560
ggcagcagca aacccaatca gtggacacta tgacagatca aaatcattga aacagaatat 1620
aaatttgtca gctcccatca tgtcccgatt cgatctcttc tttatccttg tggatgaatg 1680
taatgaggtt acagattatg ccattgccag gcgcatagta gatttgcatt caagaattga 1740
ggaatcaatt gatcgtgtct attccctcga tgatatcaga agatatcttc tctttgcaag 1800
acagtttaaa cccaagattt ccaaagagtc agaggacttc attgtggagc aatataaaca 1860
tctccgccag agagatggtt ctggagtgac caagtcttca tggaggatta cagtgcgaca 1920
gcttgagagc atgattcgtc tctctgaagc tatggctcgg atgcactgct gtgatgaggt 1980
ccaacctaaa catgtgaagg aagctttccg gttactgaat aaatcaatca tccgtgtgga 2040
aacacctgat gtcaatctag atcaagagga agagatccag atggaggtag atgagggtgc 2100
tggtggcatc aatggtcatg ctgacagccc tgctcctgtg aacgggatca atggctacaa 2160
tgaagacata aatcaagagt ctgctcccaa agcctcctta aggctgggct ctctgagta 2220
ctgccgaatc tctaacctta ttgtgcttca cctcagaaag gtggaagaag aagaggacga 2280
gtcagcatta aagaggagcg agcttgttaa ctggtacttg aaggaaatcg aatcagagat 2340
agactctgaa gaagaactta taaataaaaa aagaatcata gagaaagtta ttcatcgact 2400
cacacactat gatcatgttc taattgagct cacccaggct ggattgaaag ctccacaga 2460
gggaagtgag agctatgaag aagatcccta cttggtagtt aaccctaact acttgctcga 2520
agattgagat agtgaaagta actgaccaga gctgaggaac tgtggcacag cacctcgtgg 2580
cctggagcct ggctggagct ctgctaggga cagaagtgtt tctggaagtg atgcttccag 2640
gatttgtttt cagaaacaag aattgagttg atggtcctat gtgtcacatt catcacaggt 2700
ttcataccaa cacaggcttc agcacttcct ttggtgtgtt tcctgtccca gtgaagttgg 2760
aaccaaataa tgtgtagtct ctataaccaa tacctttgtt ttcatgtgta agaaaaggcc 2820
cattactttt aaggtatgtg ctgtcctatt gagcaaataa ctttttttca attgccagct 2880
actgctttta ttcatcaaaa taaaataact tgttctg   2917

<210> 129
<211> 821
<212> PRT
<213> Homo sapiens

<400> 129

Met	Asp	Leu	Ala	Ala	Ala	Ala	Glu	Pro	Gly	Ala	Gly	Ser	Gln	His	Leu
1				5					10					15	

Glu Val Arg Asp Glu Val Ala Glu Lys Cys Gln Lys Leu Phe Leu Asp
            20   25   30

Phe Leu Glu Glu Phe Gln Ser Ser Asp Gly Glu Ile Lys Tyr Leu Gln
        35   40   45

Leu Ala Glu Glu Leu Ile Arg Pro Glu Arg Asn Thr Leu Val Val Ser
    50   55   60

Phe Val Asp Leu Glu Gln Phe Asn Gln Gln Leu Ser Thr Thr Ile Gln
65   70   75   80

Glu Glu Phe Tyr Arg Val Tyr Pro Tyr Leu Cys Arg Ala Leu Lys Thr
                85   90   95

Phe Val Lys Asp Arg Lys Glu Ile Pro Leu Ala Lys Asp Phe Tyr Val
            100   105   110

Ala Phe Gln Asp Leu Pro Thr Arg His Lys Ile Arg Glu Leu Thr Ser
        115   120   125

Ser Arg Ile Gly Leu Leu Thr Arg Ile Ser Gly Gln Val Val Arg Thr
    130   135   140

His Pro Val His Pro Glu Leu Val Ser Gly Thr Phe Leu Cys Leu Asp
145   150   155   160

Cys Gln Thr Val Ile Arg Asp Val Glu Gln Gln Phe Lys Tyr Thr Gln
                165   170   175

Pro Asn Ile Cys Arg Asn Pro Val Cys Ala Asn Arg Arg Arg Phe Leu
            180   185   190

Leu Asp Thr Asn Lys Ser Arg Phe Val Asp Phe Gln Lys Val Arg Ile
        195   200   205

Gln Glu Thr Gln Ala Glu Leu Pro Arg Gly Ser Ile Pro Arg Ser Leu
    210   215   220

Glu Val Ile Leu Arg Ala Glu Ala Val Glu Ser Ala Gln Ala Gly Asp
225   230   235   240

Lys Cys Asp Phe Thr Gly Thr Leu Ile Val Val Pro Asp Val Ser Lys
            245   250   255

Leu Ser Thr Pro Gly Ala Arg Ala Glu Thr Asn Ser Arg Val Ser Gly
            260   265   270

Val Asp Gly Tyr Glu Thr Glu Gly Ile Arg Gly Leu Arg Ala Leu Gly
        275   280   285

Val Arg Asp Leu Ser Tyr Arg Leu Val Phe Leu Ala Cys Cys Val Ala
        290   295   300

Pro Thr Asn Pro Arg Phe Gly Gly Lys Glu Leu Arg Asp Glu Glu Gln
305   310   315   320

Thr Ala Glu Ser Ile Lys Asn Gln Met Thr Val Lys Glu Trp Glu Lys
            325   330   335

Val Phe Glu Met Ser Gln Asp Lys Asn Leu Tyr His Asn Leu Cys Thr
            340   345   350

Ser Leu Phe Pro Thr Ile His Gly Asn Asp Glu Val Lys Arg Gly Val
        355   360   365

Leu Leu Met Leu Phe Gly Gly Val Pro Lys Thr Thr Gly Glu Gly Thr
    370   375   380

Ser Leu Arg Gly Asp Ile Asn Val Cys Ile Val Gly Asp Pro Ser Thr
385   390   395   400

Ala Lys Ser Gln Phe Leu Lys His Val Glu Glu Phe Ser Pro Arg Ala
            405   410   415

```
Val Tyr Thr Ser Gly Lys Ala Ser Ser Ala Ala Gly Leu Thr Ala Ala
            420             425             430
Val Val Arg Asp Glu Glu Ser His Glu Phe Val Ile Glu Ala Gly Ala
            435             440             445
Leu Met Leu Ala Asp Asn Gly Val Cys Cys Ile Asp Glu Phe Asp Lys
        450             455             460
Met Asp Val Arg Asp Gln Val Ala Ile His Glu Ala Met Glu Gln Gln
465             470             475             480
Thr Ile Ser Ile Thr Lys Ala Gly Val Lys Ala Thr Leu Asn Ala Arg
        `       485             490             495
Thr Ser Ile Leu Ala Ala Ala Asn Pro Ile Ser Gly His Tyr Asp Arg
            500             505             510
Ser Lys Ser Leu Lys Gln Asn Ile Asn Leu Ser Ala Pro Ile Met Ser
        515             520             525
Arg Phe Asp Leu Phe Phe Ile Leu Val Asp Glu Cys Asn Glu Val Thr
    530             535             540
Asp Tyr Ala Ile Ala Arg Arg Ile Val Asp Leu His Ser Arg Ile Glu
545             550             555             560
Glu Ser Ile Asp Arg Val Tyr Ser Leu Asp Asp Ile Arg Arg Tyr Leu
            565             570             575
Leu Phe Ala Arg Gln Phe Lys Pro Lys Ile Ser Lys Glu Ser Glu Asp
            580             585             590
Phe Ile Val Glu Gln Tyr Lys His Leu Arg Gln Arg Asp Gly Ser Gly
            595             600             605
Val Thr Lys Ser Ser Trp Arg Ile Thr Val Arg Gln Leu Glu Ser Met
        610             615             620
Ile Arg Leu Ser Glu Ala Met Ala Arg Met His Cys Cys Asp Glu Val
625             630             635             640
Gln Pro Lys His Val Lys Glu Ala Phe Arg Leu Leu Asn Lys Ser Ile
            645             650             655
Ile Arg Val Glu Thr Pro Asp Val Asn Leu Asp Gln Glu Glu Glu Ile
            660             665             670
Gln Met Glu Val Asp Glu Gly Ala Gly Gly Ile Asn Gly His Ala Asp
        675             680             685
Ser Pro Ala Pro Val Asn Gly Ile Asn Gly Tyr Asn Glu Asp Ile Asn
        690             695             700
Gln Glu Ser Ala Pro Lys Ala Ser Leu Arg Leu Gly Phe Ser Glu Tyr
705             710             715             720
Cys Arg Ile Ser Asn Leu Ile Val Leu His Leu Arg Lys Val Glu Glu
            725             730             735
Glu Glu Asp Glu Ser Ala Leu Lys Arg Ser Glu Leu Val Asn Trp Tyr
            740             745             750
Leu Lys Glu Ile Glu Ser Glu Ile Asp Ser Glu Glu Glu Leu Ile Asn
        755             760             765
Lys Lys Arg Ile Ile Glu Lys Val Ile His Arg Leu Thr His Tyr Asp
        770             775             780
His Val Leu Ile Glu Leu Thr Gln Ala Gly Leu Lys Gly Ser Thr Glu
785             790             795             800
Gly Ser Glu Ser Tyr Glu Glu Asp Pro Tyr Leu Val Val Asn Pro Asn
            805             810             815
Tyr Leu Leu Glu Asp
            820
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<210> 130
<211> 786
<212> DNA
<213> Homo sapiens
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195

<400> 130
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cgggcgaagc agcgcgggca gcgagatgca gcaccgaggc ttcctcctcc tcaccctcct 60
cgccctgctg gcgctcacct ccgcggtcgc caaaaagaaa gataaggtga agaagggcgg 120
cccgggggagc gagtgcgctg agtgggcctg ggggccctgc accccccagca gcaaggattg 180
cggcgtgggt ttccgcgagg gcacctgcgg ggcccagacc cagcgcatcc ggtgcagggt 240
gccctgcaac tggaagaagg agtttggagc cgactgcaag tacaagtttg agaactgggg 300
tgcgtgtgat gggggcacag gcaccaaagt ccgccaaggc accctgaaga aggcgcgcta 360
caatgctcag tgccaggaga ccatccgcgt caccaagccc tgcaccccca agaccaaagc 420
aaaggccaaa gccaagaaag ggaagggaaa ggactagacg ccaagcctgg atgccaagga 480
gcccctggtg tcacatgggg cctggccacg ccctccctct cccaggcccg agatgtgacc 540
caccagtgcc ttctgtctgc tcgttagctt taatcaatca tgccctgcct tgtccctctc 600
actccccagc cccacccccta agtgcccaaa gtggggaggg acaagggatt ctgggaagct 660
tgagcctccc ccaaagcaat gtgagtccca gagcccgctt ttgttcttcc ccacaattcc 720
attactaaga aacacatcaa ataaactgac tttttccccc caataaaagc tcttcttttt 780
taatat                                                                 786
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<210> 131
<211> 143
<212> PRT
<213> Homo sapiens

<400> 131
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Met Gln His Arg Gly Phe Leu Leu Leu Thr Leu Leu Ala Leu Leu Ala
1               5                   10                  15
Leu Thr Ser Ala Val Ala Lys Lys Lys Asp Lys Val Lys Lys Gly Gly
            20                  25                  30
Pro Gly Ser Glu Cys Ala Glu Trp Ala Trp Gly Pro Cys Thr Pro Ser
        35                  40                  45
Ser Lys Asp Cys Gly Val Gly Phe Arg Glu Gly Thr Cys Gly Ala Gln
    50                  55                  60
Thr Gln Arg Ile Arg Cys Arg Val Pro Cys Asn Trp Lys Lys Glu Phe
65                  70                  75                  80
Gly Ala Asp Cys Lys Tyr Lys Phe Glu Asn Trp Gly Ala Cys Asp Gly
                85                  90                  95
Gly Thr Gly Thr Lys Val Arg Gln Gly Thr Leu Lys Lys Ala Arg Tyr
            100                 105                 110
Asn Ala Gln Cys Gln Glu Thr Ile Arg Val Thr Lys Pro Cys Thr Pro
            115                 120                 125
Lys Thr Lys Ala Lys Ala Lys Ala Lys Lys Gly Lys Gly Lys Asp
    130                 135                 140
```

<210> 132
<211> 603
<212> DNA
<213> Homo sapiens

<400> 132
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cgtgctgcta cacaagaacc ctgagactga cctgcaggac gaaaccatga gagcctgat 60
ccttcttgcc atcctggccg ccttagcggt agtaactttg tgttatgaat cacatgaaag 120
catggaatct tatgaactta atcccttcat taacaggaga aatgcaaata ccttcatatc 180
ccctcagcag agatggagag ctaaagtcca agagaggatc cgagaacgct ctaagcctgt 240
ccacgagctc aatagggaag cctgtgatga ctacagactt tgcgaacgct acgccatggt 300
ttatggatac aatgctgcct ataatcgcta cttcaggaag cgccgagggg ccaaatgaga 360
ctgagggaag aaaaaaaatc tcttttttc tggaggctgg cacctgattt tgtatccccc 420
tgtagcagca ttactgaaat acataggctt atatacaatg cttctttcct gtatattctc 480
ttgtctggct gcaccccttt ttcccgcccc cagattgata agtaatgaaa gtgcactgca 540
gtgagggtca aaggagagtc aacatatgtg attgttccat aataaacttc tggtgtgata 600
ctt                                                                    603
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<210> 133
<211> 103
<212> PRT
<213> Homo sapiens

<400> 133
Met Lys Ser Leu Ile Leu Leu Ala Ile Leu Ala Ala Leu Ala Val Val
1   5   10   15
Thr Leu Cys Tyr Glu Ser His Glu Ser Met Glu Ser Tyr Glu Leu Asn
            20   25   30
Pro Phe Ile Asn Arg Arg Asn Ala Asn Thr Phe Ile Ser Pro Gln Gln
            35   40   45
Arg Trp Arg Ala Lys Val Gln Glu Arg Ile Arg Glu Arg Ser Lys Pro
    50   55   60
Val His Glu Leu Asn Arg Glu Ala Cys Asp Asp Tyr Arg Leu Cys Glu
65   70   75   80
Arg Tyr Ala Met Val Tyr Gly Tyr Asn Ala Ala Tyr Asn Arg Tyr Phe
                85   90   95
Arg Lys Arg Arg Gly Ala Lys
            100

<210> 134
<211> 1778
<212> DNA
<213> Homo sapiens

<400> 134
tagaagttta caatgaagtt tcttctaata ctgctcctgc aggccactgc ttctggagct 60
cttcccctga acagctctac aagcctggaa aaaataatg tgctatttgg tgagagatac 120
ttagaaaaat tttatggcct tgagataaac aaacttccag tgacaaaaat gaaatatagt 180
ggaaacttaa tgaaggaaaa aatccaagaa atgcagcact tcttgggtct gaaagtgacc 240
gggcaactgg acacatctac cctggagatg atgcacgcac ctcgatgtgg agtccccgat 300
ctccatcatt tcagggaaat gccaggggggg cccgtatgga ggaaacatta tatcacctac 360
agaatcaata attacacacc tgacatgaac cgtgaggatg ttgactacgc aatccggaaa 420
gctttccaag tatggagtaa tgttaccccc ttgaaattca gcaagattaa cacaggcatg 480
gctgacattt tggtggtttt tgcccgtgga gctcatggag acttccatgc ttttgatggc 540
aaaggtggaa tcctagccca tgcttttgga cctggatctg gcattggagg ggatgcacat 600
ttcgatgagg acgaattctg gactacacat tcaggaggca caaacttgtt cctcactgct 660
gttcacgaga ttggccattc cttaggtctt ggccattcta gtgatccaaa ggctgtaatg 720
ttccccacct acaaatatgt cgacatcaac acatttcgcc tctctgctga tgacatacgt 780
ggcattcagt ccctgtatgg agacccaaaa gagaaccaac gcttgccaaa tcctgacaat 840
tcagaaccag ctctctgtga ccccaatttg agttttgatg ctgtcactac cgtgggaaat 900
aagatctttt tcttcaaaga caggttcttc tggctgaagg tttctgagag accaaagacc 960
agtgttaatt taatttcttc cttatggcca accttgccat ctggcattga agctgcttat 1020
gaaattgaag ccagaaatca agtttttctt tttaaagatg acaaatactg gttaattagc 1080
aatttaagac cagagccaaa ttatcccaag agcatacatt cttttggttt tcctaacttt 1140
gtgaaaaaaa ttgatgcagc tgtttttaac ccacgttttt ataggaccta cttctttgta 1200
gataaccagt attggaggta tgatgaaagg agacagatga tggaccctgg ttatcccaaa 1260
ctgattacca agaacttcca aggaatcggg cctaaaattg atgcagtctt ctattctaaa 1320
aacaaatact actatttctt ccaaggatct aaccaatttg aatatgactt cctactccaa 1380
cgtatcacca aaacactgaa aagcaatagc tggtttggtt gttagaaatg gtgtaattaa 1440
tggttttttgt tagttcactt cagcttaata agtatttatt gcatatttgc tatgtcctca 1500
gtgtaccact acttagagat atgtatcata aaaataaaat ctgtaaacca taggtaatga 1560
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taaaattaag tatatatatt ttggctcaaa taaaattg 1778

```
<210> 135
<211> 470
<212> PRT
<213> Homo sapiens

<400> 135
Met Lys Phe Leu Leu Ile Leu Leu Leu Gln Ala Thr Ala Ser Gly Ala
1               5                   10                  15
Leu Pro Leu Asn Ser Ser Thr Ser Leu Glu Lys Asn Asn Val Leu Phe
                20                  25                  30
Gly Glu Arg Tyr Leu Glu Lys Phe Tyr Gly Leu Glu Ile Asn Lys Leu
            35                  40                  45
Pro Val Thr Lys Met Lys Tyr Ser Gly Asn Leu Met Lys Glu Lys Ile
        50                  55                  60
Gln Glu Met Gln His Phe Leu Gly Leu Lys Val Thr Gly Gln Leu Asp
65                  70                  75                  80
Thr Ser Thr Leu Glu Met Met His Ala Pro Arg Cys Gly Val Pro Asp
                85                  90                  95
Leu His His Phe Arg Glu Met Pro Gly Gly Pro Val Trp Arg Lys His
            100                 105                 110
Tyr Ile Thr Tyr Arg Ile Asn Asn Tyr Thr Pro Asp Met Asn Arg Glu
        115                 120                 125
Asp Val Asp Tyr Ala Ile Arg Lys Ala Phe Gln Val Trp Ser Asn Val
    130                 135                 140
Thr Pro Leu Lys Phe Ser Lys Ile Asn Thr Gly Met Ala Asp Ile Leu
145                 150                 155                 160
Val Val Phe Ala Arg Gly Ala His Gly Asp Phe His Ala Phe Asp Gly
                165                 170                 175
Lys Gly Gly Ile Leu Ala His Ala Phe Gly Pro Gly Ser Gly Ile Gly
            180                 185                 190
Gly Asp Ala His Phe Asp Glu Asp Glu Phe Trp Thr Thr His Ser Gly
        195                 200                 205
Gly Thr Asn Leu Phe Leu Thr Ala Val His Glu Ile Gly His Ser Leu
    210                 215                 220
Gly Leu Gly His Ser Ser Asp Pro Lys Ala Val Met Phe Pro Thr Tyr
225                 230                 235                 240
Lys Tyr Val Asp Ile Asn Thr Phe Arg Leu Ser Ala Asp Asp Ile Arg
                245                 250                 255
Gly Ile Gln Ser Leu Tyr Gly Asp Pro Lys Glu Asn Gln Arg Leu Pro
            260                 265                 270
Asn Pro Asp Asn Ser Glu Pro Ala Leu Cys Asp Pro Asn Leu Ser Phe
        275                 280                 285
Asp Ala Val Thr Thr Val Gly Asn Lys Ile Phe Phe Phe Lys Asp Arg
    290                 295                 300
Phe Phe Trp Leu Lys Val Ser Glu Arg Pro Lys Thr Ser Val Asn Leu
305                 310                 315                 320
Ile Ser Ser Leu Trp Pro Thr Leu Pro Ser Gly Ile Glu Ala Ala Tyr
                325                 330                 335
Glu Ile Glu Ala Arg Asn Gln Val Phe Leu Phe Lys Asp Asp Lys Tyr
            340                 345                 350
Trp Leu Ile Ser Asn Leu Arg Pro Glu Pro Asn Tyr Pro Lys Ser Ile
        355                 360                 365
His Ser Phe Gly Phe Pro Asn Phe Val Lys Lys Ile Asp Ala Ala Val
    370                 375                 380
Phe Asn Pro Arg Phe Tyr Arg Thr Tyr Phe Phe Val Asp Asn Gln Tyr
385                 390                 395                 400
Trp Arg Tyr Asp Glu Arg Arg Gln Met Met Asp Pro Gly Tyr Pro Lys
                405                 410                 415
```

```
Leu Ile Thr Lys Asn Phe Gln Gly Ile Gly Pro Lys Ile Asp Ala Val
            420                 425             430
Phe Tyr Ser Lys Asn Lys Tyr Tyr Tyr Phe Phe Gln Gly Ser Asn Gln
            435             440             445
Phe Glu Tyr Asp Phe Leu Leu Gln Arg Ile Thr Lys Thr Leu Lys Ser
    450                 455             460
Asn Ser Trp Phe Gly Cys
465             470
```

```
<210> 136
<211> 1821
<212> DNA
<213> Homo sapiens

<400> 136
acaaggaggc aggcaagaca gcaaggcata gagacaacat agagctaagt aaagccagtg 60
gaaatgaaga gtcttccaat cctactgttg ctgtgcgtgg cagtttgctc agcctatcca 120
ttggatggag ctgcaagggg tgaggacacc agcatgaacc ttgttcagaa atatctagaa 180
aactactacg acctcaaaaa agatgtgaaa cagtttgtta ggagaaagga cagtggtcct 240
gttgttaaaa aaatccgaga aatgcagaag ttccttggat tggaggtgac ggggaagctg 300
gactccgaca ctctggaggt gatgcgcaag cccaggtgtg gagttcctga tgttggtcac 360
ttcagaacct ttcctggcat cccgaagtgg aggaaacccc accttacata caggattgtg 420
aattatacac agatttgcc aaaagatgct gttgattctg ctgttgagaa agctctgaaa 480
gtctgggaag aggtgactcc actcacattc tccaggctgt atgaaggaga ggctgatata 540
atgatctctt ttgcagttag agaacatgga gactttacc cttttgatgg acctggaaat 600
gttttggccc atgcctatgc ccctgggcca gggattaatg gagatgccca ctttgatgat 660
gatgaacaat ggacaaagga tacaacaggg accaatttat ttctcgttgc tgctcatgaa 720
attggccact ccctgggtct ctttcactca gccaacactg aagctttgat gtacccactc 780
tatcactcac tcacagacct gactcggttc cgcctgtctc aagatgatat aaatggcatt 840
cagtccctct atggacctcc ccctgactcc cctgagaccc ccctggtacc cacggaacct 900
gtccctccag aacctgggac gccagccaac tgtgatcctg ctttgtcctt tgatgctgtc 960
agcactctga ggggagaaat cctgatcttt aaagacaggc acttttggcg caaatccctc 1020
aggaagcttg aacctgaatt gcatttgatc tcttcatttt ggccatctct tccttcaggc 1080
gtggatgccg catatgaagt tactagcaag gacctcgttt tcattttaa aggaaatcaa 1140
ttctgggcca tcagaggaaa tgaggtacga gctggatacc aagaggcat ccacacccta 1200
ggtttccctc caaccgtgag gaaaatcgat gcagccattt ctgataagga aagaacaaa 1260
acatatttct ttgtagagga caaatactgg agatttgatg agaagagaaa ttccatggag 1320
ccaggctttc ccaagcaaat agctgaagac tttccaggga ttgactcaaa gattgatgct 1380
gttttttgaag aatttgggtt ctttttatttc tttactggat cttcacagtt ggagtttgac 1440
ccaaatgcaa agaaagtgac acacactttg aagagtaaca gctggcttaa ttgttgaaag 1500
agatatgtag aaggcacaat atgggcactt taaatgaagc taataattct tcacctaagt 1560
ctctgtgaat tgaaatgttc gttttctcct gcctgtgctg tgactcgagt cacactcaag 1620
ggaacttgag cgtgaatctg tatcttgccg gtcattttta tgttattaca gggcattcaa 1680
atgggctgct gcttagcttg caccttgtca catagagtga tctttcccaa gagaagggga 1740
agcactcgtg tgcaacagac aagtgactgt atctgtgtag actatttgct tatttaataa 1800
agacgatttg tcagttgttt t                                          1821
```

```
<210> 137
<211> 477
<212> PRT
<213> Homo sapiens

<400> 137
Met Lys Ser Leu Pro Ile Leu Leu Leu Leu Cys Val Ala Val Cys Ser
1               5                   10                  15
Ala Tyr Pro Leu Asp Gly Ala Ala Arg Gly Glu Asp Thr Ser Met Asn
            20                  25                  30
Leu Val Gln Lys Tyr Leu Glu Asn Tyr Tyr Asp Leu Lys Lys Asp Val
```

```
              35                    40                    45
Lys Gln Phe Val Arg Arg Lys Asp Ser Gly Pro Val Val Lys Lys Ile
    50                    55                    60
Arg Glu Met Gln Lys Phe Leu Gly Leu Glu Val Thr Gly Lys Leu Asp
65                    70                    75                    80
Ser Asp Thr Leu Glu Val Met Arg Lys Pro Arg Cys Gly Val Pro Asp
                  85                    90                    95
Val Gly His Phe Arg Thr Phe Pro Gly Ile Pro Lys Trp Arg Lys Thr
              100                   105                   110
His Leu Thr Tyr Arg Ile Val Asn Tyr Thr Pro Asp Leu Pro Lys Asp
              115                   120                   125
Ala Val Asp Ser Ala Val Glu Lys Ala Leu Lys Val Trp Glu Glu Val
          130                   135                   140
Thr Pro Leu Thr Phe Ser Arg Leu Tyr Glu Gly Glu Ala Asp Ile Met
145                   150                   155                   160
Ile Ser Phe Ala Val Arg Glu His Gly Asp Phe Tyr Pro Phe Asp Gly
                  165                   170                   175
Pro Gly Asn Val Leu Ala His Ala Tyr Ala Pro Gly Pro Gly Ile Asn
              180                   185                   190
Gly Asp Ala His Phe Asp Asp Asp Glu Gln Trp Thr Lys Asp Thr Thr
          195                   200                   205
Gly Thr Asn Leu Phe Leu Val Ala Ala His Glu Ile Gly His Ser Leu
      210                   215                   220
Gly Leu Phe His Ser Ala Asn Thr Glu Ala Leu Met Tyr Pro Leu Tyr
225                   230                   235                   240
His Ser Leu Thr Asp Leu Thr Arg Phe Arg Leu Ser Gln Asp Asp Ile
                  245                   250                   255
Asn Gly Ile Gln Ser Leu Tyr Gly Pro Pro Pro Asp Ser Pro Glu Thr
              260                   265                   270
Pro Leu Val Pro Thr Glu Pro Val Pro Pro Glu Pro Gly Thr Pro Ala
          275                   280                   285
Asn Cys Asp Pro Ala Leu Ser Phe Asp Ala Val Ser Thr Leu Arg Gly
      290                   295                   300
Glu Ile Leu Ile Phe Lys Asp Arg His Phe Trp Arg Lys Ser Leu Arg
305                   310                   315                   320
Lys Leu Glu Pro Glu Leu His Leu Ile Ser Ser Phe Trp Pro Ser Leu
                  325                   330                   335
Pro Ser Gly Val Asp Ala Ala Tyr Glu Val Thr Ser Lys Asp Leu Val
              340                   345                   350
Phe Ile Phe Lys Gly Asn Gln Phe Trp Ala Ile Arg Gly Asn Glu Val
          355                   360                   365
Arg Ala Gly Tyr Pro Arg Gly Ile His Thr Leu Gly Phe Pro Pro Thr
      370                   375                   380
Val Arg Lys Ile Asp Ala Ala Ile Ser Asp Lys Glu Lys Asn Lys Thr
385                   390                   395                   400
Tyr Phe Phe Val Glu Asp Lys Tyr Trp Arg Phe Asp Glu Lys Arg Asn
                  405                   410                   415
Ser Met Glu Pro Gly Phe Pro Lys Gln Ile Ala Glu Asp Phe Pro Gly
              420                   425                   430
Ile Asp Ser Lys Ile Asp Ala Val Phe Glu Glu Phe Gly Phe Phe Tyr
          435                   440                   445
Phe Phe Thr Gly Ser Ser Gln Leu Glu Phe Asp Pro Asn Ala Lys Lys
      450                   455                   460
Val Thr His Thr Leu Lys Ser Asn Ser Trp Leu Asn Cys
465                   470                   475
```

<210> 138
<211> 1127

```
<212> DNA
<213> Homo sapiens

<400> 138
accaaatcaa ccataggtcc aagaacaatt gtctctggac ggcagctatg cgactcaccg 60
tgctgtgtgc tgtgtgcctg ctgcctggca gcctggccct gccgctgcct caggaggcgg 120
gaggcatgag tgagctacag tgggaacagg ctcaggacta tctcaagaga ttttatctct 180
atgactcaga aacaaaaaat gccaacagtt tagaagccaa actcaaggag atgcaaaaat 240
tctttggcct acctataact ggaatgttaa actcccgcgt catagaaata atgcagaagc 300
ccagatgtgg agtgccagat gttgcagaat actcactatt tccaaatagc ccaaatgga 360
cttccaaagt ggtcacctac aggatcgtat catatactcg agacttaccg catattacag 420
tggatcgatt agtgtcaaag gctttaaaca tgtgggggcaa agagatcccc ctgcatttca 480
ggaaagttgt atggggaact gctgacatca tgattggctt gcgcgagga gctcatgggg 540
actcctaccc atttgatggg ccaggaaaca cgctggctca tgcctttgcg cctgggacag 600
gtctcggagg agatgctcac ttcgatgagg atgaacgctg acggatggt agcagtctag 660
ggattaactt cctgtatgct gcaactcatg aacttggcca ttcctttgggt atgggacatt 720
cctctgatcc taatgcagtg atgtatccaa cctatggaaa tggagatccc caaaatttta 780
aactttccca ggatgatatt aaaggcattc agaaactata tggaaagaga agtaattcaa 840
gaaagaaata gaaacttcag gcagaacatc cattcattca ttcattggat tgtatatcat 900
tgttgcacaa tcagaattga taagcactgt cctccactc catttagcaa ttatgtcacc 960
cttttttatt gcagttggtt tttgaatgtc tttcactcct tttattggtt aaactccttt 1020
atggtgtgac tgtgtcttat tccatctatg agctttgtca gtgcgcgtag atgtcaataa 1080
atgttacata cacaaataaa taaaatgttt attccatggt aaattta      1127
```

```
<210> 139
<211> 267
<212> PRT
<213> Homo sapiens

<400> 139
Met Arg Leu Thr Val Leu Cys Ala Val Cys Leu Leu Pro Gly Ser Leu
1               5                   10                  15
Ala Leu Pro Leu Pro Gln Glu Ala Gly Gly Met Ser Glu Leu Gln Trp
            20                  25                  30
Glu Gln Ala Gln Asp Tyr Leu Lys Arg Phe Tyr Leu Tyr Asp Ser Glu
        35                  40                  45
Thr Lys Asn Ala Asn Ser Leu Glu Ala Lys Leu Lys Glu Met Gln Lys
    50                  55                  60
Phe Phe Gly Leu Pro Ile Thr Gly Met Leu Asn Ser Arg Val Ile Glu
65                  70                  75                  80
Ile Met Gln Lys Pro Arg Cys Gly Val Pro Asp Val Ala Glu Tyr Ser
                85                  90                  95
Leu Phe Pro Asn Ser Pro Lys Trp Thr Ser Lys Val Val Thr Tyr Arg
            100                 105                 110
Ile Val Ser Tyr Thr Arg Asp Leu Pro His Ile Thr Val Asp Arg Leu
        115                 120                 125
Val Ser Lys Ala Leu Asn Met Trp Gly Lys Glu Ile Pro Leu His Phe
    130                 135                 140
Arg Lys Val Val Trp Gly Thr Ala Asp Ile Met Ile Gly Phe Ala Arg
145                 150                 155                 160
Gly Ala His Gly Asp Ser Tyr Pro Phe Asp Gly Pro Gly Asn Thr Leu
                165                 170                 175
Ala His Ala Phe Ala Pro Gly Thr Gly Leu Gly Gly Asp Ala His Phe
            180                 185                 190
Asp Glu Asp Glu Arg Trp Thr Asp Gly Ser Ser Leu Gly Ile Asn Phe
        195                 200                 205
Leu Tyr Ala Ala Thr His Glu Leu Gly His Ser Leu Gly Met Gly His
    210                 215                 220
Ser Ser Asp Pro Asn Ala Val Met Tyr Pro Thr Tyr Gly Asn Gly Asp
```

225   230   235   240
Pro Gln Asn Phe Lys Leu Ser Gln Asp Asp Ile Lys Gly Ile Gln Lys
                    245   250   255
Leu Tyr Gly Lys Arg Ser Asn Ser Arg Lys Lys
              260   265

<210> 140
<211> 1078
<212> DNA
<213> Homo sapiens

<400> 140
aagaacaatt gtctctggac ggcagctatg cgactcaccg tgctgtgtgc tgtgtgcctg   60
ctgcctggca gcctggccct gccgctgcct caggaggcgg gaggcatgag tgagctacag 120
tgggaacagg ctcaggacta tctcaagaga tttttatctct atgactcaga aacaaaaaat 180
gccaacagtt tagaagccaa actcaaggag atgcaaaaat tctttggcct acctataact 240
ggaatgttaa actcccgcgt catagaaata atgcagaagc ccagatgtgg agtgccagat 300
gttgcagaat actcactatt tccaaatagc ccaaaatgga cttccaaagt ggtcacctac 360
aggatcgtat catatactcg agacttaccg catattacag tggatcgatt agtgtcaaag 420
gctttaaaca tgtggggcaa agagatcccc ctgcatttca ggaaagttgt atggggaact 480
gctgacatca tgattggctt tgcgcgagga gctcatgggg actcctaccc atttgatggg 540
ccaggaaaca cgctggctca tgcctttgcg cctgggacag gtctcggagg agatgctcac 600
ttcgatgagg atgaacgctg gacggatggt agcagtctag ggattaactt cctgtatgct 660
gcaactcatg aacttggcca ttctttgggt atgggacatt cctctgatcc taatgcagtg 720
atgtatccaa cctatggaaa tggagatccc caaaatttta aactttccca ggatgatatt 780
aaaggcattc agaaactata tggaaagaga agtaattcaa gaaagaaata gaaacttcag 840
gcagaacatc cattcattca ttcattggat tgtatatcat gttgcacaa tcagaattga   900
taagcactgt cctccactc catttagcaa ttatgtcacc cttttttatt gcagttggtt   960
tttgaatgtc tttcactcct tttattggtt aaactccttt atggtgtgac tgtgtcttat 1020
tccatctatg agctttgtca gtgcgcgtag atgtcaataa atgttacata cacaaata   1078

<210> 141
<211> 2334
<212> DNA
<213> Homo sapiens

<400> 141
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gggctgctgc tttgctgccc ccagacagcg ccagtccacc cttgtgctct tccctggaga 120
cctgagaacc aatctcaccg acaggcagct ggcagaggaa tacctgtacc gctatggtta 180
cactcgggtg gcagagatgc gtggagagtc gaaatctctg gggcctgcgc tgctgcttct 240
ccagaagcaa ctgtccctgc ccgagaccgg tgagctggat agcgccacgc tgaaggccat 300
gcgaacccca cggtgcgggg tcccagacct gggcagattc caaaccttg agggcgacct 360
caagtggcac caccacaaca tcacctattg gatccaaaac tactcggaag acttgccgcg 420
ggcggtgatt gacgacgcct ttgcccgcgc cttcgcactg tggagcgcgg tgacgccgct 480
caccttcact cgcgtgtaca gccgggacgc agacatcgtc atccagtttg tgtcgcgga   540
gcacggagac gggtatccct tcgacgggaa ggacgggctc ctggcacacg cctttcctcc 600
tggccccggc attcagggag acgcccattt cgacgatgac gagttgtggt ccctgggcaa 660
gggcgtcgtg gttccaactc ggtttggaaa cgcagatggc gcggcctgcc acttccccctt 720
catcttcgag ggccgctcct actctgcctg caccaccgac ggtcgctccg acggcttgcc 780
ctggtgcagt accacggcca actacgacac cgacgaccgg tttggcttct gccccagcga 840
gagactctac acccgggacg gcaatgctga tgggaaaccc tgccagtttc cattcatctt 900
ccaaggccaa tcctactccg cctgcaccac ggacggtcgc tccgacggct accgctggtg 960
cgccaccacc gccaactacg accgggacaa gctcttcggc ttctgcccga cccgagctga 1020
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gggtaaggag tactcgacct gtaccagcga gggccgcgga tgggcgcc tctggtgcgc 1140
taccacctcg aactttgaca gcgacaagaa gtggggcttc tgcccggacc aaggatacag 1200
tttgttcctc gtggcggcgc atgagttcgg ccacgcgctg ggcttagatc attcctcagt 1260

```
gccggaggcg ctcatgtacc ctatgtaccg cttcactgag gggccccccct tgcataagga 1320
cgacgtgaat ggcatccggc acctctatgg tcctcgccct gaacctgagc cacggcctcc 1380
aaccaccacc acaccgcagc ccacggctcc cccgacggtc tgccccaccg gaccccccac 1440
tgtccacccc tcagagcgcc ccacagctgg ccccacaggt cccccctcag ctggccccac 1500
aggtcccccc actgctggcc cttctacggc cactactgtg cctttgagtc cggtggacga 1560
tgcctgcaac gtgaacatct tcgacgccat cgcggagatt gggaaccagc tgtatttgtt 1620
caaggatggg aagtactggc gattctctga gggcaggggg agccggccgc agggcccctt 1680
ccttatcgcc gacaagtggc ccgcgctgcc ccgcaagctg gactcggtct ttgaggagcc 1740
gctctccaag aagcttttct tcttctctgg gcgccaggtg tgggtgtaca caggcgcgtc 1800
ggtgctgggc ccgaggcgtc tggacaagct gggcctggga gccgacgtgg cccaggtgac 1860
cggggccctc cggagtggca gggggaagat gctgctgttc agcgggcggc gcctctggag 1920
gttcgacgtg aaggcgcaga tggtggatcc ccggagcgcc agcgaggtgg accggatgtt 1980
ccccgggggtg cctttggaca cgcacgacgt cttccagtac cgagagaaag cctatttctg 2040
ccaggaccgc ttctactggc gcgtgagttc ccggagtgag ttgaaccagg tggaccaagt 2100
gggctacgtg acctatgaca tcctgcagtg ccctgaggac tagggctccc gtcctgcttt 2160
gcagtgccat gtaaatcccc actgggacca accctgggga aggagccagt ttgccggata 2220
caaactggta ttctgttctg gaggaaaggg aggagtggag gtgggctggg ccctctcttc 2280
tcacctttgt tttttgttgg agtgtttcta ataaacttgg attctctaac cttt      2334
```

<210> 142
<211> 707
<212> PRT
<213> Homo sapiens

<400> 142

```
Met Ser Leu Trp Gln Pro Leu Val Leu Val Leu Leu Val Leu Gly Cys
1               5                   10                  15
Cys Phe Ala Ala Pro Arg Gln Arg Gln Ser Thr Leu Val Leu Phe Pro
                20                  25                  30
Gly Asp Leu Arg Thr Asn Leu Thr Asp Arg Gln Leu Ala Glu Glu Tyr
        35                  40                  45
Leu Tyr Arg Tyr Gly Tyr Thr Arg Val Ala Glu Met Arg Gly Glu Ser
    50                  55                  60
Lys Ser Leu Gly Pro Ala Leu Leu Leu Leu Gln Lys Gln Leu Ser Leu
65                  70                  75                  80
Pro Glu Thr Gly Glu Leu Asp Ser Ala Thr Leu Lys Ala Met Arg Thr
                85                  90                  95
Pro Arg Cys Gly Val Pro Asp Leu Gly Arg Phe Gln Thr Phe Glu Gly
            100                 105                 110
Asp Leu Lys Trp His His His Asn Ile Thr Tyr Trp Ile Gln Asn Tyr
        115                 120                 125
Ser Glu Asp Leu Pro Arg Ala Val Ile Asp Asp Ala Phe Ala Arg Ala
    130                 135                 140
Phe Ala Leu Trp Ser Ala Val Thr Pro Leu Thr Phe Thr Arg Val Tyr
145                 150                 155                 160
Ser Arg Asp Ala Asp Ile Val Ile Gln Phe Gly Val Ala Glu His Gly
                165                 170                 175
Asp Gly Tyr Pro Phe Asp Gly Lys Asp Gly Leu Leu Ala His Ala Phe
            180                 185                 190
Pro Pro Gly Pro Gly Ile Gln Gly Asp Ala His Phe Asp Asp Asp Glu
        195                 200                 205
Leu Trp Ser Leu Gly Lys Gly Val Val Val Pro Thr Arg Phe Gly Asn
    210                 215                 220
Ala Asp Gly Ala Ala Cys His Phe Pro Phe Ile Phe Glu Gly Arg Ser
225                 230                 235                 240
Tyr Ser Ala Cys Thr Thr Asp Gly Arg Ser Asp Gly Leu Pro Trp Cys
                245                 250                 255
Ser Thr Thr Ala Asn Tyr Asp Thr Asp Asp Arg Phe Gly Phe Cys Pro
            260                 265                 270
```

```
Ser Glu Arg Leu Tyr Thr Arg Asp Gly Asn Ala Asp Gly Lys Pro Cys
        275             280             285
Gln Phe Pro Phe Ile Phe Gln Gly Gln Ser Tyr Ser Ala Cys Thr Thr
    290             295             300
Asp Gly Arg Ser Asp Gly Tyr Arg Trp Cys Ala Thr Thr Ala Asn Tyr
305             310             315             320
Asp Arg Asp Lys Leu Phe Gly Phe Cys Pro Thr Arg Ala Asp Ser Thr
            325             330             335
Val Met Gly Gly Asn Ser Ala Gly Glu Leu Cys Val Phe Pro Phe Thr
            340             345             350
Phe Leu Gly Lys Glu Tyr Ser Thr Cys Thr Ser Glu Gly Arg Gly Asp
            355             360             365
Gly Arg Leu Trp Cys Ala Thr Thr Ser Asn Phe Asp Ser Asp Lys Lys
    370             375             380
Trp Gly Phe Cys Pro Asp Gln Gly Tyr Ser Leu Phe Leu Val Ala Ala
385             390             395             400
His Glu Phe Gly His Ala Leu Gly Leu Asp His Ser Ser Val Pro Glu
            405             410             415
Ala Leu Met Tyr Pro Met Tyr Arg Phe Thr Glu Gly Pro Pro Leu His
            420             425             430
Lys Asp Asp Val Asn Gly Ile Arg His Leu Tyr Gly Pro Arg Pro Glu
            435             440             445
Pro Glu Pro Arg Pro Pro Thr Thr Thr Thr Pro Gln Pro Thr Ala Pro
    450             455             460
Pro Thr Val Cys Pro Thr Gly Pro Pro Thr Val His Pro Ser Glu Arg
465             470             475             480
Pro Thr Ala Gly Pro Thr Gly Pro Pro Ser Ala Gly Pro Thr Gly Pro
            485             490             495
Pro Thr Ala Gly Pro Ser Thr Ala Thr Thr Val Pro Leu Ser Pro Val
            500             505             510
Asp Asp Ala Cys Asn Val Asn Ile Phe Asp Ala Ile Ala Glu Ile Gly
            515             520             525
Asn Gln Leu Tyr Leu Phe Lys Asp Gly Lys Tyr Trp Arg Phe Ser Glu
    530             535             540
Gly Arg Gly Ser Arg Pro Gln Gly Pro Phe Leu Ile Ala Asp Lys Trp
545             550             555             560
Pro Ala Leu Pro Arg Lys Leu Asp Ser Val Phe Glu Glu Pro Leu Ser
            565             570             575
Lys Lys Leu Phe Phe Phe Ser Gly Arg Gln Val Trp Val Tyr Thr Gly
            580             585             590
Ala Ser Val Leu Gly Pro Arg Arg Leu Asp Lys Leu Gly Leu Gly Ala
            595             600             605
Asp Val Ala Gln Val Thr Gly Ala Leu Arg Ser Gly Arg Gly Lys Met
    610             615             620
Leu Leu Phe Ser Gly Arg Arg Leu Trp Arg Phe Asp Val Lys Ala Gln
625             630             635             640
Met Val Asp Pro Arg Ser Ala Ser Glu Val Asp Arg Met Phe Pro Gly
            645             650             655
Val Pro Leu Asp Thr His Asp Val Phe Gln Tyr Arg Glu Lys Ala Tyr
            660             665             670
Phe Cys Gln Asp Arg Phe Tyr Trp Arg Val Ser Ser Arg Ser Glu Leu
            675             680             685
Asn Gln Val Asp Gln Val Gly Tyr Val Thr Tyr Asp Ile Leu Gln Cys
    690             695             700
Pro Glu Asp
705
```

<210> 143

<211> 2217
<212> DNA
<213> Homo sapiens

<400> 143
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ggccggccac tcccgtctgc tgtgacgcgc ggacagagag ctaccggtgg acccacggtg   60
cctccctccc tgggatctac acagaccatg gccttgccaa cggctcgacc cctgttgggg  120
tcctgtggga cccccgccct cggcagcctc ctgttcctgc tcttcagcct cggatgggtg  180
cagccctcga ggaccctggc tggagagaca gggcaggagg ctgcacccct ggacggagtc  240
ctggccaacc cacctaacat ttccagcctc tccctcgcc aactccttgg cttcccgtgt  300
gcggaggtgt ccggcctgag cacggagcgt gtccgggagc tggctgtggc cttggcacag  360
aagaatgtca agctctcaac agagcagctg cgctgtctgg ctcaccggct ctctgagccc  420
cccgaggacc tggacgccct cccattggac ctgctgctat tcctcaaccc agatgcgttc  480
tcggggcccc aggcctgcac ccgtttcttc tcccgcatca cgaaggccaa tgtggacctg  540
ctcccgaggg gggctcccga gcgacagcgg ctgctgcctg cggctctggc ctgctggggt  600
gtgcggggggt ctctgctgag cgaggctgat gtgcgggctc tgggaggcct ggcttgcgac  660
ctgcctgggc gctttgtggc cgagtcggcc gaagtgctgc tacccccggct ggtgagctgc  720
ccgggacccc tggaccagga ccagcaggag gcagccaggg cggctctgca gggcggggga  780
cccccctacg gcccccgtc gacatggtct gtctccacga tggacgctct gcggggcctg  840
ctgccgtgc tgggccagcc catcatccgc agcatcccgc agggcatcgt ggccgcgtgg  900
cggcaacgct cctctcggga cccatcctgg cggcagcctg aacggaccat cctccggccg  960
cggttccggc gggaagtgga gaagacagcc tgtccttcag gcaagaaggc ccgcgagata 1020
gacgagagcc tcatcttcta caagaagtgg gagctggaag cctgcgtgga tgcggccctg 1080
ctggccaccc agatggaccg cgtgaacgcc atcccctta cctacgagca gctggacgtc 1140
ctaaagcata aactggatga gctctaccca caaggttacc ccgagtctgt gatccagcac 1200
ctgggctacc tcttcctcaa gatgagccct gaggacattc gcaagtggaa tgtgacgtcc 1260
ctggagaccc tgaaggcttt gcttgaagtc aacaaagggc acgaaatgag tcctcaggct 1320
cctcggcggc ccctcccaca ggtggccacc ctgatcgacc gctttgtgaa gggaaggggc 1380
cagctagaca aagacaccct agacaccctg accgccttct accctgggta cctgtgctcc 1440
ctcagccccg aggagctgag ctccgtgccc cccagcagca tctgggcggt caggcccccag 1500
gacctggaca cgtgtgaccc aaggcagctg gacgtcctct atcccaaggc ccgccttgct 1560
ttccagaaca tgaacgggtc cgaatacttc gtgaagatcc agtccttcct gggtgggggc 1620
cccacggagg atttgaaggc gctcagtcag cagaatgtga gcatggactt ggccacgttc 1680
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gggggcagag ctgggggcgt ggaggtgggc gctctgagtc accctctct ctgtagaggc 1980
cctctcgggg acgccctgcc tcctaggacc tggacctgtt ctcaccgtcc tggcactgct 2040
cctagcctcc accctggcct gagggccca ctcccttgct ggccccagcc ctgctgggga 2100
tccccgcctg gccaggagca ggcacgggtg atccccgttc cacccaaga gaactcgcgc 2160
tcagtaaacg ggaacatgcc ccctgcagac acgtaaaaaa aaaaaaaaa aaaaaa        2217
```

<210> 144
<211> 702
<212> PRT
<213> Homo sapiens

<400> 144
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Met Ala Leu Pro Thr Ala Arg Pro Leu Leu Gly Ser Cys Gly Thr Pro
1               5                   10                  15
Ala Leu Gly Ser Leu Leu Phe Leu Leu Phe Ser Leu Gly Trp Val Gln
                20                  25                  30
Pro Ser Arg Thr Leu Ala Gly Glu Thr Gly Gln Glu Ala Ala Pro Leu
            35                  40                  45
Asp Gly Val Leu Ala Asn Pro Pro Asn Ile Ser Ser Leu Ser Pro Arg
        50                  55                  60
Gln Leu Leu Gly Phe Pro Cys Ala Glu Val Ser Gly Leu Ser Thr Glu
65                  70                  75                  80
```

```
Arg Val Arg Glu Leu Ala Val Ala Leu Ala Gln Lys Asn Val Lys Leu
              85                  90                  95
Ser Thr Glu Gln Leu Arg Cys Leu Ala His Arg Leu Ser Glu Pro Pro
              100                 105                 110
Glu Asp Leu Asp Ala Leu Pro Leu Asp Leu Leu Leu Phe Leu Asn Pro
              115                 120                 125
Asp Ala Phe Ser Gly Pro Gln Ala Cys Thr Arg Phe Phe Ser Arg Ile
    130                 135                 140
Thr Lys Ala Asn Val Asp Leu Leu Pro Arg Gly Ala Pro Glu Arg Gln
145                 150                 155                 160
Arg Leu Leu Pro Ala Ala Leu Ala Cys Trp Gly Val Arg Gly Ser Leu
              165                 170                 175
Leu Ser Glu Ala Asp Val Arg Ala Leu Gly Gly Leu Ala Cys Asp Leu
              180                 185                 190
Pro Gly Arg Phe Val Ala Glu Ser Ala Glu Val Leu Leu Pro Arg Leu
              195                 200                 205
Val Ser Cys Pro Gly Pro Leu Asp Gln Asp Gln Gln Glu Ala Ala Arg
    210                 215                 220
Ala Ala Leu Gln Gly Gly Gly Pro Pro Tyr Gly Pro Pro Ser Thr Trp
225                 230                 235                 240
Ser Val Ser Thr Met Asp Ala Leu Arg Gly Leu Leu Pro Val Leu Gly
              245                 250                 255
Gln Pro Ile Ile Arg Ser Ile Pro Gln Gly Ile Val Ala Ala Trp Arg
              260                 265                 270
Gln Arg Ser Ser Arg Asp Pro Ser Trp Arg Gln Pro Glu Arg Thr Ile
              275                 280                 285
Leu Arg Pro Arg Phe Arg Arg Glu Val Glu Lys Thr Ala Cys Pro Ser
    290                 295                 300
Gly Lys Lys Ala Arg Glu Ile Asp Glu Ser Leu Ile Phe Tyr Lys Lys
305                 310                 315                 320
Trp Glu Leu Glu Ala Cys Val Asp Ala Ala Leu Leu Ala Thr Gln Met
              325                 330                 335
Asp Arg Val Asn Ala Ile Pro Phe Thr Tyr Glu Gln Leu Asp Val Leu
              340                 345                 350
Lys His Lys Leu Asp Glu Leu Tyr Pro Gln Gly Tyr Pro Glu Ser Val
              355                 360                 365
Ile Gln His Leu Gly Tyr Leu Phe Leu Lys Met Ser Pro Glu Asp Ile
    370                 375                 380
Arg Lys Trp Asn Val Thr Ser Leu Glu Thr Leu Lys Ala Leu Leu Glu
385                 390                 395                 400
Val Asn Lys Gly His Glu Met Ser Pro Gln Ala Pro Arg Arg Pro Leu
              405                 410                 415
Pro Gln Val Ala Thr Leu Ile Asp Arg Phe Val Lys Gly Arg Gly Gln
              420                 425                 430
Leu Asp Lys Asp Thr Leu Asp Thr Leu Thr Ala Phe Tyr Pro Gly Tyr
              435                 440                 445
Leu Cys Ser Leu Ser Pro Glu Glu Leu Ser Ser Val Pro Pro Ser Ser
    450                 455                 460
Ile Trp Ala Val Arg Pro Gln Asp Leu Asp Thr Cys Asp Pro Arg Gln
465                 470                 475                 480
Leu Asp Val Leu Tyr Pro Lys Ala Arg Leu Ala Phe Gln Asn Met Asn
              485                 490                 495
Gly Ser Glu Tyr Phe Val Lys Ile Gln Ser Phe Leu Gly Gly Ala Pro
              500                 505                 510
Thr Glu Asp Leu Lys Ala Leu Ser Gln Gln Asn Val Ser Met Asp Leu
              515                 520                 525
Ala Thr Phe Met Lys Leu Arg Thr Asp Ala Val Leu Pro Leu Thr Val
    530                 535                 540
Ala Glu Val Gln Lys Leu Leu Gly Pro His Val Glu Gly Leu Lys Ala
```

```
545                    550                    555                    560
Glu Glu Arg His Arg Pro Val Arg Asp Trp Ile Leu Arg Gln Arg Gln
            565                    570                    575
Asp Asp Leu Asp Thr Leu Gly Leu Gly Leu Gln Gly Gly Ile Pro Asn
            580                    585                    590
Gly Tyr Leu Val Leu Asp Leu Ser Val Gln Gly Gly Arg Gly Gly Gln
            595                    600                    605
Ala Arg Ala Gly Gly Arg Ala Gly Gly Val Glu Val Gly Ala Leu Ser
    610                    615                    620
His Pro Ser Leu Cys Arg Gly Pro Leu Gly Asp Ala Leu Pro Pro Arg
625                    630                    635                    640
Thr Trp Thr Cys Ser His Arg Pro Gly Thr Ala Pro Ser Leu His Pro
                645                    650                    655
Gly Leu Arg Ala Pro Leu Pro Cys Trp Pro Gln Pro Cys Trp Gly Ser
            660                    665                    670
Pro Pro Gly Gln Glu Gln Ala Arg Val Ile Pro Val Pro Pro Gln Glu
        675                    680                    685
Asn Ser Arg Ser Val Asn Gly Asn Met Pro Pro Ala Asp Thr
        690                    695                    700
```

```
<210> 145
<211> 2135
<212> DNA
<213> Homo sapiens

<400> 145
ggccggccac tcccgtctgc tgtgacgcgc ggacagagag ctaccggtgg acccacggtg   60
cctccctccc tgggatctac acagaccatg gccttgccaa cggctcgacc cctgttggggg  120
tcctgtggga cccccgccct cggcagcctc ctgttcctgc tcttcagcct cggatgggtg  180
cagccctcga ggaccctggc tggagagaca gggcaggagg ctgcacccct ggacggagtc  240
ctggccaacc cacctaacat ttccctcgcc aactccttgg cttcccgtgt  300
gcggaggtgt ccggcctgag cacggagcgt gtccgggagc tggctgtgtg cttggcacag  360
aagaatgtca agctctcaac agagcagctg cgctgtctgg ctcaccggct ctctgagccc  420
cccgaggacc tggacgccct cccattggac ctgctgctat tcctcaaccc agatgcgttc  480
tcggggcccc aggcctgcac ccgtttcttc tcccgcatca cgaaggccaa tgtggacctg  540
ctcccgaggg gggctcccga gcgacagcgg ctgctgcctg cggctctggc ctgctggggt  600
gtgcggggggt ctctgctgag cgaggctgat gtgcgggctc tgggaggcct ggcttgcgac  660
ctgcctgggc gctttgtggc cgagtcggcc gaagtgctgc taccccggct ggtgagctgc  720
ccgggacccc tggaccagga ccagcaggag cagccaggg cggctctgca gggcggggga  780
ccccctacg gccccccgtc gacatggtct gtctccacga tggacgctct gcggggcctg  840
ctgcccgtgc tgggccagcc catcatccgc agcatcccgc agggcatcgt ggccgcgtgg  900
cggcaacgct cctctcggga cccatcctgg cggcagcctg aacggaccat cctccggccg  960
cggttccggc gggaagtgga gaagacagcc tgtccttcag gcaagaaggc ccgcgagata  1020
gacgagagcc tcatcttcta caagaagtgg gagctggaag cctgcgtgga tgcggccctg  1080
ctggccaccc agatggaccg cgtgaacgcc atccccttca cctacgagca gctggacgtc  1140
ctaaagcata aactggatga gctctaccca caaggttacc ccgagtctgt gatccagcac  1200
ctgggctacc tcttcctcaa gatgagccct gaggacattc gcaagtggaa tgtgacgtcc  1260
ctggagaccc tgaaggcttt gcttgaagtc aacaaagggc acgaaatgag tcctcaggct  1320
cctcggcggc ccctcccaca ggtggccacc ctgatcgacc gctttgtgaa gggaaggggc  1380
cagctagaca agacaccct agacaccctg accgccttct accctgggta cctgtgctcc  1440
ctcagccccg aggagctgag ctccgtgccc cccagcagca tctgggcggt caggccccag  1500
gacctggaca cgtgtgaccc aaggcagctg gacgtcctct atcccaaggc ccgccttgct  1560
ttccagaaca tgaacgggtc cgaatacttc gtgaagatcc agtccttcct gggtgggggcc  1620
cccacggagg atttgaaggc gctcagtcag cagaatgtga gcatggactt ggccacgttc  1680
atgaagctgc ggacggatgc ggtgctgccg ttgactgtgg ctgaggtgca gaaacttctg  1740
ggaccccacg tggagggcct gaaggcggag gagcggcacc gcccggtgcg ggactggatc  1800
ctacggcagc ggcaggacga cctggacacg ctggggctgg gctacagggg cggcatcccc  1860
aacggctacc tggtcctaga cctcagcgtg caagaggccc tctcggggac gccctgcctc  1920
```

```
ctaggacctg gacctgttct caccgtcctg gcactgctcc tagcctccac cctggcctga 1980
gggcccccact cccttgctgg ccccagccct gctggggatc cccgcctggc caggagcagg 2040
cacgggtgat ccccgttcca ccccaagaga actcgcgctc agtaaacggg aacatgcccc 2100
ctgcagacac gtaaaaaaaa aaaaaaaaaa aaaaa                              2135
```

<210> 146
<211> 630
<212> PRT
<213> Homo sapiens

<400> 146

Met	Ala	Leu	Pro	Thr	Ala	Arg	Pro	Leu	Leu	Gly	Ser	Cys	Gly	Thr	Pro
1				5					10					15	
Ala	Leu	Gly	Ser	Leu	Leu	Phe	Leu	Leu	Phe	Ser	Leu	Gly	Trp	Val	Gln
			20					25					30		
Pro	Ser	Arg	Thr	Leu	Ala	Gly	Glu	Thr	Gly	Gln	Glu	Ala	Ala	Pro	Leu
			35				40					45			
Asp	Gly	Val	Leu	Ala	Asn	Pro	Pro	Asn	Ile	Ser	Ser	Leu	Ser	Pro	Arg
		50				55					60				
Gln	Leu	Leu	Gly	Phe	Pro	Cys	Ala	Glu	Val	Ser	Gly	Leu	Ser	Thr	Glu
65					70					75					80
Arg	Val	Arg	Glu	Leu	Ala	Val	Ala	Leu	Ala	Gln	Lys	Asn	Val	Lys	Leu
				85					90					95	
Ser	Thr	Glu	Gln	Leu	Arg	Cys	Leu	Ala	His	Arg	Leu	Ser	Glu	Pro	Pro
			100					105					110		
Glu	Asp	Leu	Asp	Ala	Leu	Pro	Leu	Asp	Leu	Leu	Leu	Phe	Leu	Asn	Pro
			115				120					125			
Asp	Ala	Phe	Ser	Gly	Pro	Gln	Ala	Cys	Thr	Arg	Phe	Phe	Ser	Arg	Ile
		130				135					140				
Thr	Lys	Ala	Asn	Val	Asp	Leu	Leu	Pro	Arg	Gly	Ala	Pro	Glu	Arg	Gln
145					150					155					160
Arg	Leu	Leu	Pro	Ala	Ala	Leu	Ala	Cys	Trp	Gly	Val	Arg	Gly	Ser	Leu
				165					170					175	
Leu	Ser	Glu	Ala	Asp	Val	Arg	Ala	Leu	Gly	Gly	Leu	Ala	Cys	Asp	Leu
			180					185					190		
Pro	Gly	Arg	Phe	Val	Ala	Glu	Ser	Ala	Glu	Val	Leu	Leu	Pro	Arg	Leu
		195				200					205				
Val	Ser	Cys	Pro	Gly	Pro	Leu	Asp	Gln	Asp	Gln	Gln	Glu	Ala	Ala	Arg
	210				215					220					
Ala	Ala	Leu	Gln	Gly	Gly	Gly	Pro	Pro	Tyr	Gly	Pro	Pro	Ser	Thr	Trp
225				230				235					240		
Ser	Val	Ser	Thr	Met	Asp	Ala	Leu	Arg	Gly	Leu	Leu	Pro	Val	Leu	Gly
			245				250					255			
Gln	Pro	Ile	Ile	Arg	Ser	Ile	Pro	Gln	Gly	Ile	Val	Ala	Ala	Trp	Arg
			260				265					270			
Gln	Arg	Ser	Ser	Arg	Asp	Pro	Ser	Trp	Arg	Gln	Pro	Glu	Arg	Thr	Ile
		275				280					285				
Leu	Arg	Pro	Arg	Phe	Arg	Arg	Glu	Val	Glu	Lys	Thr	Ala	Cys	Pro	Ser
	290				295					300					
Gly	Lys	Lys	Ala	Arg	Glu	Ile	Asp	Glu	Ser	Leu	Ile	Phe	Tyr	Lys	Lys
305				310				315					320		
Trp	Glu	Leu	Glu	Ala	Cys	Val	Asp	Ala	Ala	Leu	Leu	Ala	Thr	Gln	Met
			325				330					335			
Asp	Arg	Val	Asn	Ala	Ile	Pro	Phe	Thr	Tyr	Glu	Gln	Leu	Asp	Val	Leu
		340				345					350				
Lys	His	Lys	Leu	Asp	Glu	Leu	Tyr	Pro	Gln	Gly	Tyr	Pro	Glu	Ser	Val
	355				360					365					
Ile	Gln	His	Leu	Gly	Tyr	Leu	Phe	Leu	Lys	Met	Ser	Pro	Glu	Asp	Ile
370				375				380							

```
Arg Lys Trp Asn Val Thr Ser Leu Glu Thr Leu Lys Ala Leu Leu Glu
385                     390             395                 400
Val Asn Lys Gly His Glu Met Ser Pro Gln Ala Pro Arg Arg Pro Leu
            405                     410             415
Pro Gln Val Ala Thr Leu Ile Asp Arg Phe Val Lys Gly Arg Gly Gln
            420                 425             430
Leu Asp Lys Asp Thr Leu Asp Thr Leu Thr Ala Phe Tyr Pro Gly Tyr
        435             440             445
Leu Cys Ser Leu Ser Pro Glu Glu Leu Ser Ser Val Pro Pro Ser Ser
    450             455             460
Ile Trp Ala Val Arg Pro Gln Asp Leu Asp Thr Cys Asp Pro Arg Gln
465             470             475             480
Leu Asp Val Leu Tyr Pro Lys Ala Arg Leu Ala Phe Gln Asn Met Asn
            485             490             495
Gly Ser Glu Tyr Phe Val Lys Ile Gln Ser Phe Leu Gly Gly Ala Pro
        500             505             510
Thr Glu Asp Leu Lys Ala Leu Ser Gln Gln Asn Val Ser Met Asp Leu
    515             520             525
Ala Thr Phe Met Lys Leu Arg Thr Asp Ala Val Leu Pro Leu Thr Val
    530             535             540
Ala Glu Val Gln Lys Leu Leu Gly Pro His Val Glu Gly Leu Lys Ala
545             550             555                 560
Glu Glu Arg His Arg Pro Val Arg Asp Trp Ile Leu Arg Gln Arg Gln
            565             570             575
Asp Asp Leu Asp Thr Leu Gly Leu Gly Leu Gln Gly Gly Ile Pro Asn
        580             585             590
Gly Tyr Leu Val Leu Asp Leu Ser Val Gln Glu Ala Leu Ser Gly Thr
        595             600             605
Pro Cys Leu Leu Gly Pro Gly Pro Val Leu Thr Val Leu Ala Leu Leu
    610             615             620
Leu Ala Ser Thr Leu Ala
625             630
```

```
<210> 147
<211> 2105
<212> DNA
<213> Homo sapiens

<400> 147
ggccggccac tcccgtctgc tgtgacgcgc ggacagagag ctaccggtgg acccacggtg 60
cctccctccc tgggatctac acagaccatg gccttgccaa cggctcgacc cctgttggg 120
tcctgtggga ccccgcccct cggcagcctc ctgttcctgc tcttcagcct cggatgggtg 180
cagccctcga ggaccctggc tggagagaca gggcaggagg ctgcacccct ggacggagtc 240
ctggccaacc cacctaacat ttccagcctc tccctcgcc aactccttgg cttcccgtgt 300
gcggaggtgt ccggcctgag cacggagcgt gtccgggagc tggctgtggc cttggcacag 360
aagaatgtca agctctcaac agagcagctg cgctgtctgg ctcaccggct ctctgagccc 420
cccgaggacc tggacgccct cccattggac ctgctgctat tcctcaaccc agatgcgttc 480
tcggggcccc aggcctgcac ccgtttcttc tcccgcatca cgaaggccaa tgtggacctg 540
ctcccgaggg gggctcccga gcgacagcgg ctgctgcctg cggctctggc ctgctggggt 600
gtgcgggggt ctctgctgag cgaggctgat gtgcgggctc tgggaggcct ggcttgcgac 660
ctgcctgggc gctttgtggc cgagtcggcc gaagtgctgc taccccggct ggtgagctgc 720
ccgggacccc tggaccagga ccagcaggag cagccaggg cggctctgca gggcgggga 780
ccccctacg gcccccgtc gacatggtct gtctccacga tggacgctct gcgggggcctg 840
ctgcccgtgc tgggccagcc catcatccgc agcatccgc agggcatcgt ggccgcgtgg 900
cggcaacgct cctctcggga cccatcctgg cggcagcctg aacggaccat cctccggccg 960
cggttccggc gggaagtgga gaagacagcc tgtccttcag gcaagaaggc ccgcgagata 1020
gacgagagcc tcatcttcta caagaagtgg gagctggaag cctgcgtgga tgcggccctg 1080
ctggccaccc agatggaccg cgtgaacgcc atccccttca cctacgagca gctggacgtc 1140
```

```
ctaaagcata aactggatga gctctaccca caaggttacc ccgagtctgt gatccagcac 1200
ctgggctacc tcttcctcaa gatgagccct gaggacattc gcaagtggaa tgtgacgtcc 1260
ctggagaccc tgaaggcttt gcttgaagtc aacaaagggc acgaaatgag tcctcaggct 1320
cctcggcggc ccctcccaca ggtggccacc ctgatcgacc gctttgtgaa gggaaggggc 1380
cagctagaca aagacaccct agacaccctg accgccttct accctgggta cctgtgctcc 1440
ctcagccccg aggagctgag ctccgtgccc cccagcagca tctgggcggt caggccccag 1500
gacctggaca cgtgtgaccc aaggcagctg gacgtcctct atcccaaggc ccgccttgct 1560
ttccagaaca tgaacgggtc cgaatacttc gtgaagatcc agtccttcct gggtgggggcc 1620
cccacggagg atttgaaggc gctcagtcag cagaatgtga gcatggactt ggccacgttc 1680
atgaagctgc ggacggatgc ggtgctgccg ttgactgtgg ctgaggtgca gaaacttctg 1740
ggaccccacg tggagggcct gaaggcggag gagcggcacc gcccggtgcg ggactggatc 1800
ctacggcagc ggcaggacga cctggacacg ctggggctgg ggctacaggg cggcatcccc 1860
aacggctacc tggtcctaga cctcagcgtg caaggacctg gacctgttct caccgtcctg 1920
gcactgctcc tagcctccac cctggcctga gggccccact cccttgctgg ccccagccct 1980
gctggggatc cccgcctggc caggagcagg cacgggtgat ccccgttcca ccccaagaga 2040
actcgcgctc agtaaacggg aacatgcccc ctgcagacac gtaaaaaaaa aaaaaaaaaa 2100
aaaaa                                                                2105
```

```
<210> 148
<211> 620
<212> PRT
<213> Homo sapiens
```

```
<400> 148
Met Ala Leu Pro Thr Ala Arg Pro Leu Leu Gly Ser Cys Gly Thr Pro
1               5                   10                  15
Ala Leu Gly Ser Leu Leu Phe Leu Leu Phe Ser Leu Gly Trp Val Gln
            20                  25                  30
Pro Ser Arg Thr Leu Ala Gly Glu Thr Gly Gln Glu Ala Ala Pro Leu
        35                  40                  45
Asp Gly Val Leu Ala Asn Pro Pro Asn Ile Ser Ser Leu Ser Pro Arg
    50                  55                  60
Gln Leu Leu Gly Phe Pro Cys Ala Glu Val Ser Gly Leu Ser Thr Glu
65                  70                  75                  80
Arg Val Arg Glu Leu Ala Val Ala Leu Ala Gln Lys Asn Val Lys Leu
                85                  90                  95
Ser Thr Glu Gln Leu Arg Cys Leu Ala His Arg Leu Ser Glu Pro Pro
            100                 105                 110
Glu Asp Leu Asp Ala Leu Pro Leu Asp Leu Leu Leu Phe Leu Asn Pro
        115                 120                 125
Asp Ala Phe Ser Gly Pro Gln Ala Cys Thr Arg Phe Phe Ser Arg Ile
    130                 135                 140
Thr Lys Ala Asn Val Asp Leu Leu Pro Arg Gly Ala Pro Glu Arg Gln
145                 150                 155                 160
Arg Leu Leu Pro Ala Ala Leu Ala Cys Trp Gly Val Arg Gly Ser Leu
                165                 170                 175
Leu Ser Glu Ala Asp Val Arg Ala Leu Gly Gly Leu Ala Cys Asp Leu
            180                 185                 190
Pro Gly Arg Phe Val Ala Glu Ser Ala Glu Val Leu Leu Pro Arg Leu
        195                 200                 205
Val Ser Cys Pro Gly Pro Leu Asp Gln Asp Gln Gln Glu Ala Ala Arg
    210                 215                 220
Ala Ala Leu Gln Gly Gly Gly Pro Pro Tyr Gly Pro Pro Ser Thr Trp
225                 230                 235                 240
Ser Val Ser Thr Met Asp Ala Leu Arg Gly Leu Leu Pro Val Leu Gly
                245                 250                 255
Gln Pro Ile Ile Arg Ser Ile Pro Gln Gly Ile Val Ala Ala Trp Arg
            260                 265                 270
Gln Arg Ser Ser Arg Asp Pro Ser Trp Arg Gln Pro Glu Arg Thr Ile
```

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                275                    280                    285
Leu Arg Pro Arg Phe Arg Arg Glu Val Glu Lys Thr Ala Cys Pro Ser
    290                 295                 300
Gly Lys Lys Ala Arg Glu Ile Asp Glu Ser Leu Ile Phe Tyr Lys Lys
305                 310                 315                 320
Trp Glu Leu Glu Ala Cys Val Asp Ala Ala Leu Leu Ala Thr Gln Met
                325                 330                 335
Asp Arg Val Asn Ala Ile Pro Phe Thr Tyr Glu Gln Leu Asp Val Leu
                340                 345                 350
Lys His Lys Leu Asp Glu Leu Tyr Pro Gln Gly Tyr Pro Glu Ser Val
            355                 360                 365
Ile Gln His Leu Gly Tyr Leu Phe Leu Lys Met Ser Pro Glu Asp Ile
    370                 375                 380
Arg Lys Trp Asn Val Thr Ser Leu Glu Thr Leu Lys Ala Leu Leu Glu
385                 390                 395                 400
Val Asn Lys Gly His Glu Met Ser Pro Gln Ala Pro Arg Arg Pro Leu
                405                 410                 415
Pro Gln Val Ala Thr Leu Ile Asp Arg Phe Val Lys Gly Arg Gly Gln
                420                 425                 430
Leu Asp Lys Asp Thr Leu Asp Thr Leu Thr Ala Phe Tyr Pro Gly Tyr
            435                 440                 445
Leu Cys Ser Leu Ser Pro Glu Glu Leu Ser Ser Val Pro Pro Ser Ser
    450                 455                 460
Ile Trp Ala Val Arg Pro Gln Asp Leu Asp Thr Cys Asp Pro Arg Gln
465                 470                 475                 480
Leu Asp Val Leu Tyr Pro Lys Ala Arg Leu Ala Phe Gln Asn Met Asn
                485                 490                 495
Gly Ser Glu Tyr Phe Val Lys Ile Gln Ser Phe Leu Gly Gly Ala Pro
            500                 505                 510
Thr Glu Asp Leu Lys Ala Leu Ser Gln Gln Asn Val Ser Met Asp Leu
            515                 520                 525
Ala Thr Phe Met Lys Leu Arg Thr Asp Ala Val Leu Pro Leu Thr Val
    530                 535                 540
Ala Glu Val Gln Lys Leu Leu Gly Pro His Val Glu Gly Leu Lys Ala
545                 550                 555                 560
Glu Glu Arg His Arg Pro Val Arg Asp Trp Ile Leu Arg Gln Arg Gln
            565                 570                 575
Asp Asp Leu Asp Thr Leu Gly Leu Gly Leu Gln Gly Gly Ile Pro Asn
            580                 585                 590
Gly Tyr Leu Val Leu Asp Leu Ser Val Gln Gly Pro Gly Pro Val Leu
    595                 600                 605
Thr Val Leu Ala Leu Leu Leu Ala Ser Thr Leu Ala
    610                 615                 620
```

<210> 149
<211> 2193
<212> DNA
<213> Homo sapiens

<400> 149

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ggccggccac tcccgtctgc tgtgacgcgc ggacagagag ctaccggtgg acccacggtg 60
cctccctccc tgggatctac acagaccatg gccttgccaa cggctcgacc cctgttgggg 120
tcctgtggga cccccgccct cggcagcctc ctgttcctgc tcttcagcct cggatgggtg 180
cagccctcga ggaccctggc tggagagaca gggcaggagg ctgcacccct ggacggagtc 240
ctggccaacc cacctaacat ttccagcctc tcccctcgcc aactccttgg cttcccgtgt 300
gcggaggtgt ccggcctgag cacggagcgt gtccgggagc tggctgtggc cttggcacag 360
aagaatgtca agctctcaac agagcagctg cgctgtctgg ctcaccggct ctctgagccc 420
cccgaggacc tggacgccct cccattggac ctgctgctat tcctcaaccc agatgcgttc 480
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tcggggcccc aggcctgcac ccgtttcttc tcccgcatca cgaaggccaa tgtggacctg 540
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gtgcggggggt ctctgctgag cgaggctgat gtgcgggctc tgggaggcct ggcttgcgac 660
ctgcctgggc gctttgtggc cgagtcggcc gaagtgctgc taccccggct ggtgagctgc 720
ccgggacccc tggaccagga ccagcaggag gcagccaggg cggctctgca gggcgggggga 780
ccccctacg gcccccccgtc gacatggtct gtctccacga tggacgctct cgggggcctg 840
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cggcaacgct cctctcggga cccatcctgg cggcagcctg aacggaccat cctccggccg 960
cggttccggc gggaagtgga gaagacagcc tgtccttcag gcaagaaggc ccgcgagata 1020
gacgagagcc tcatcttcta caagaagtgg gagctggaag cctgcgtgga tgcggccctg 1080
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gacacgctgg ggctggggct acaggcggc atccccaacg ctacctggt cctagacctc 1860
agcgtgcaag gtgggcgggg cggccaggcc agggctgggg gcagagctgg gggcgtggag 1920
gtgggcgctc tgagtcaccc ctctctctgt agaggccctc tcggggacgc cctgcctcct 1980
aggacctgga cctgttctca ccgtcctggc actgctccta gcctccaccc tggcctgagg 2040
gccccactcc cttgctggcc ccagccctgc tggggatccc cgcctggcca ggagcaggca 2100
cgggtgatcc ccgttccacc ccaagagaac tcgcgctcag taaacgggaa catgccccct 2160
gcagacacgt aaaaaaaaaa aaaaaaaaaa aaa                               2193
```

```
<210> 150
<211> 694
<212> PRT
<213> Homo sapiens

<400> 150
Met Ala Leu Pro Thr Ala Arg Pro Leu Leu Gly Ser Cys Gly Thr Pro
1               5                   10                  15
Ala Leu Gly Ser Leu Leu Phe Leu Leu Phe Ser Leu Gly Trp Val Gln
            20                  25                  30
Pro Ser Arg Thr Leu Ala Gly Glu Thr Gly Gln Glu Ala Ala Pro Leu
            35                  40                  45
Asp Gly Val Leu Ala Asn Pro Pro Asn Ile Ser Ser Leu Ser Pro Arg
        50                  55                  60
Gln Leu Leu Gly Phe Pro Cys Ala Glu Val Ser Gly Leu Ser Thr Glu
65                  70                  75                  80
Arg Val Arg Glu Leu Ala Val Ala Leu Ala Gln Lys Asn Val Lys Leu
                85                  90                  95
Ser Thr Glu Gln Leu Arg Cys Leu Ala His Arg Leu Ser Glu Pro Pro
            100                 105                 110
Glu Asp Leu Asp Ala Leu Pro Leu Asp Leu Leu Leu Phe Leu Asn Pro
            115                 120                 125
Asp Ala Phe Ser Gly Pro Gln Ala Cys Thr Arg Phe Phe Ser Arg Ile
        130                 135                 140
Thr Lys Ala Asn Val Asp Leu Leu Pro Arg Gly Ala Pro Glu Arg Gln
145                 150                 155                 160
Arg Leu Leu Pro Ala Ala Leu Ala Cys Trp Gly Val Arg Gly Ser Leu
                165                 170                 175
Leu Ser Glu Ala Asp Val Arg Ala Leu Gly Gly Leu Ala Cys Asp Leu
```

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                    180                 185                 190
Pro Gly Arg Phe Val Ala Glu Ser Ala Glu Val Leu Leu Pro Arg Leu
        195                 200                 205
Val Ser Cys Pro Gly Pro Leu Asp Gln Asp Gln Gln Glu Ala Ala Arg
        210                 215                 220
Ala Ala Leu Gln Gly Gly Gly Pro Pro Tyr Gly Pro Pro Ser Thr Trp
225                 230                 235                 240
Ser Val Ser Thr Met Asp Ala Leu Arg Gly Leu Leu Pro Val Leu Gly
                    245                 250                 255
Gln Pro Ile Ile Arg Ser Ile Pro Gln Gly Ile Val Ala Ala Trp Arg
                    260                 265                 270
Gln Arg Ser Ser Arg Asp Pro Ser Trp Arg Gln Pro Glu Arg Thr Ile
        275                 280                 285
Leu Arg Pro Arg Phe Arg Arg Glu Val Glu Lys Thr Ala Cys Pro Ser
        290                 295                 300
Gly Lys Lys Ala Arg Glu Ile Asp Glu Ser Leu Ile Phe Tyr Lys Lys
305                 310                 315                 320
Trp Glu Leu Glu Ala Cys Val Asp Ala Ala Leu Leu Ala Thr Gln Met
                    325                 330                 335
Asp Arg Val Asn Ala Ile Pro Phe Thr Tyr Glu Gln Leu Asp Val Leu
                    340                 345                 350
Lys His Lys Leu Asp Glu Leu Tyr Pro Gln Gly Tyr Pro Glu Ser Val
        355                 360                 365
Ile Gln His Leu Gly Tyr Leu Phe Leu Lys Met Ser Pro Glu Asp Ile
        370                 375                 380
Arg Lys Trp Asn Val Thr Ser Leu Glu Thr Leu Lys Ala Leu Leu Glu
385                 390                 395                 400
Val Asn Lys Gly His Glu Met Ser Pro Gln Val Ala Thr Leu Ile Asp
                    405                 410                 415
Arg Phe Val Lys Gly Arg Gly Gln Leu Asp Lys Asp Thr Leu Asp Thr
                    420                 425                 430
Leu Thr Ala Phe Tyr Pro Gly Tyr Leu Cys Ser Leu Ser Pro Glu Glu
                    435                 440                 445
Leu Ser Ser Val Pro Pro Ser Ser Ile Trp Ala Val Arg Pro Gln Asp
        450                 455                 460
Leu Asp Thr Cys Asp Pro Arg Gln Leu Asp Val Leu Tyr Pro Lys Ala
465                 470                 475                 480
Arg Leu Ala Phe Gln Asn Met Asn Gly Ser Glu Tyr Phe Val Lys Ile
                    485                 490                 495
Gln Ser Phe Leu Gly Gly Ala Pro Thr Glu Asp Leu Lys Ala Leu Ser
        500                 505                 510
Gln Gln Asn Val Ser Met Asp Leu Ala Thr Phe Met Lys Leu Arg Thr
        515                 520                 525
Asp Ala Val Leu Pro Leu Thr Val Ala Glu Val Gln Lys Leu Leu Gly
        530                 535                 540
Pro His Val Glu Gly Leu Lys Ala Glu Glu Arg His Arg Pro Val Arg
545                 550                 555                 560
Asp Trp Ile Leu Arg Gln Arg Gln Asp Asp Leu Asp Thr Leu Gly Leu
                    565                 570                 575
Gly Leu Gln Gly Gly Ile Pro Asn Gly Tyr Leu Val Leu Asp Leu Ser
        580                 585                 590
Val Gln Gly Gly Arg Gly Gly Gln Ala Arg Ala Gly Gly Arg Ala Gly
        595                 600                 605
Gly Val Glu Val Gly Ala Leu Ser His Pro Ser Leu Cys Arg Gly Pro
        610                 615                 620
Leu Gly Asp Ala Leu Pro Pro Arg Thr Trp Thr Cys Ser His Arg Pro
625                 630                 635                 640
Gly Thr Ala Pro Ser Leu His Pro Gly Leu Arg Ala Pro Leu Pro Cys
                .645                 650                 655
```

Trp Pro Gln Pro Cys Trp Gly Ser Pro Pro Gly Gln Glu Gln Ala Arg
        660   665   670
Val Ile Pro Val Pro Pro Gln Glu Asn Ser Arg Ser Val Asn Gly Asn
        675   680   685
Met Pro Pro Ala Asp Thr
        690

<210> 151
<211> 2081
<212> DNA
<213> Homo sapiens

<400> 151
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cctccctccc tgggatctac acagaccatg gccttgccaa cggctcgacc cctgttgggg 120
tcctgtggga cccccgccct cggcagcctc ctgttcctgc tcttcagcct cggatgggtg 180
cagccctcga ggaccctggc tggagagaca gggcaggagg ctgcacccct ggacggagtc 240
ctggccaacc cacctaacat ttccagcctc tccctcgcc aactccttgg cttcccgtgt 300
gcggaggtgt ccggcctgag cacggagcgt gtccgggagc tggctgtggc cttggcacag 360
aagaatgtca agctctcaac agagcagctg cgctgtctgg ctcaccggct ctctgagccc 420
cccgaggacc tggacgccct cccattggac ctgctgctat tcctcaaccc agatgcgttc 480
tcggggcccc aggcctgcac ccgtttcttc tcccgcatca cgaaggccaa tgtggacctg 540
ctcccgaggg gggctcccga gcgacagcgg ctgctgcctg cggctctggc ctgctggggt 600
gtgcgggggt ctctgctgag cgaggctgat gtgcgggctc tgggaggcct ggcttgcgac 660
ctgcctgggc gctttgtggc cgagtcggcc gaagtgctgc taccccggct ggtgagctgc 720
ccgggacccc tggaccagga ccagcaggag cagccagggg cggctctgca gggcggggga 780
ccccctacg gcccccgtc gacatggtct gtctccacga tggacgctct gcggggcctg 840
ctgcccgtgc tgggccagcc catcatccgc agcatcccgc agggcatcgt ggccgcgtgg 900
cggcaacgct cctctcggga cccatcctgg cggcagcctg aacggaccat cctccggccg 960
cggttccggc gggaagtgga gaagacagcc tgtccttcag gcaagaaggc ccgcgagata 1020
gacgagagcc tcatcttcta caagaagtgg gagctggaag cctgcgtgga tgcggccctg 1080
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gccaccctga tcgaccgctt tgtgaaggga agggccagc tagacaaaga caccctagac 1380
accctgaccg ccttctaccc tgggtacctg tgctccctca gccccgagga gctgagctcc 1440
gtgccccca gcagcatctg ggcggtcagg ccccaggacc tggacacgtg tgacccaagg 1500
cagctggacg tcctctatcc caaggcccgc cttgctttcc agaacatgaa cgggtccgaa 1560
tacttcgtga gatccagtc cttcctgggt ggggccccca cggaggattt gaaggcgctc 1620
agtcagcaga atgtgagcat ggacttggcc acgttcatga gctgcggac ggatgcggtg 1680
ctgccgttga ctgtggctga ggtgcagaaa cttctgggac cccacgtgga gggcctgaag 1740
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gcctgagggc cccactccct tgctggcccc agccctgctg gggatccccg cctggccagg 1980
agcaggcacg ggtgatcccc gttccacccc aagagaactc gcgctcagta aacgggaaca 2040
tgccccctgc agacacgtaa aaaaaaaaaa aaaaaaaaaa a 2081

<210> 152
<211> 612
<212> PRT
<213> Homo sapiens

<400> 152
Met Ala Leu Pro Thr Ala Arg Pro Leu Leu Gly Ser Cys Gly Thr Pro
 1   5   10   15
Ala Leu Gly Ser Leu Leu Phe Leu Leu Phe Ser Leu Gly Trp Val Gln

```
                    20                    25                    30
Pro Ser Arg Thr Leu Ala Gly Glu Thr Gly Gln Glu Ala Ala Pro Leu
        35                    40                    45
Asp Gly Val Leu Ala Asn Pro Pro Asn Ile Ser Ser Leu Ser Pro Arg
    50                    55                    60
Gln Leu Leu Gly Phe Pro Cys Ala Glu Val Ser Gly Leu Ser Thr Glu
65                    70                    75                    80
Arg Val Arg Glu Leu Ala Val Ala Leu Ala Gln Lys Asn Val Lys Leu
                85                    90                    95
Ser Thr Glu Gln Leu Arg Cys Leu Ala His Arg Leu Ser Glu Pro Pro
            100                   105                   110
Glu Asp Leu Asp Ala Leu Pro Leu Asp Leu Leu Leu Phe Leu Asn Pro
        115                   120                   125
Asp Ala Phe Ser Gly Pro Gln Ala Cys Thr Arg Phe Phe Ser Arg Ile
    130                   135                   140
Thr Lys Ala Asn Val Asp Leu Leu Pro Arg Gly Ala Pro Glu Arg Gln
145                   150                   155                   160
Arg Leu Leu Pro Ala Ala Leu Ala Cys Trp Gly Val Arg Gly Ser Leu
                165                   170                   175
Leu Ser Glu Ala Asp Val Arg Ala Leu Gly Gly Leu Ala Cys Asp Leu
            180                   185                   190
Pro Gly Arg Phe Val Ala Glu Ser Ala Glu Val Leu Leu Pro Arg Leu
        195                   200                   205
Val Ser Cys Pro Gly Pro Leu Asp Gln Asp Gln Gln Glu Ala Ala Arg
    210                   215                   220
Ala Ala Leu Gln Gly Gly Gly Pro Pro Tyr Gly Pro Pro Ser Thr Trp
225                   230                   235                   240
Ser Val Ser Thr Met Asp Ala Leu Arg Gly Leu Leu Pro Val Leu Gly
                245                   250                   255
Gln Pro Ile Ile Arg Ser Ile Pro Gln Gly Ile Val Ala Ala Trp Arg
            260                   265                   270
Gln Arg Ser Ser Arg Asp Pro Ser Trp Arg Gln Pro Glu Arg Thr Ile
        275                   280                   285
Leu Arg Pro Arg Phe Arg Arg Glu Val Glu Lys Thr Ala Cys Pro Ser
    290                   295                   300
Gly Lys Lys Ala Arg Glu Ile Asp Glu Ser Leu Ile Phe Tyr Lys Lys
305                   310                   315                   320
Trp Glu Leu Glu Ala Cys Val Asp Ala Ala Leu Leu Ala Thr Gln Met
                325                   330                   335
Asp Arg Val Asn Ala Ile Pro Phe Thr Tyr Glu Gln Leu Asp Val Leu
            340                   345                   350
Lys His Lys Leu Asp Glu Leu Tyr Pro Gln Gly Tyr Pro Glu Ser Val
        355                   360                   365
Ile Gln His Leu Gly Tyr Leu Phe Leu Lys Met Ser Pro Glu Asp Ile
    370                   375                   380
Arg Lys Trp Asn Val Thr Ser Leu Glu Thr Leu Lys Ala Leu Leu Glu
385                   390                   395                   400
Val Asn Lys Gly His Glu Met Ser Pro Gln Val Ala Thr Leu Ile Asp
                405                   410                   415
Arg Phe Val Lys Gly Arg Gly Gln Leu Asp Lys Asp Thr Leu Asp Thr
            420                   425                   430
Leu Thr Ala Phe Tyr Pro Gly Tyr Leu Cys Ser Leu Ser Pro Glu Glu
        435                   440                   445
Leu Ser Ser Val Pro Pro Ser Ser Ile Trp Ala Val Arg Pro Gln Asp
    450                   455                   460
Leu Asp Thr Cys Asp Pro Arg Gln Leu Asp Val Leu Tyr Pro Lys Ala
465                   470                   475                   480
Arg Leu Ala Phe Gln Asn Met Asn Gly Ser Glu Tyr Phe Val Lys Ile
                485                   490                   495
```

215

```
Gln Ser Phe Leu Gly Gly Ala Pro Thr Glu Asp Leu Lys Ala Leu Ser
            500                     505             510
Gln Gln Asn Val Ser Met Asp Leu Ala Thr Phe Met Lys Leu Arg Thr
            515                     520             525
Asp Ala Val Leu Pro Leu Thr Val Ala Glu Val Gln Lys Leu Leu Gly
    530                     535             540
Pro His Val Glu Gly Leu Lys Ala Glu Glu Arg His Arg Pro Val Arg
545                     550             555              560
Asp Trp Ile Leu Arg Gln Arg Gln Asp Asp Leu Asp Thr Leu Gly Leu
                565             570              575
Gly Leu Gln Gly Gly Ile Pro Asn Gly Tyr Leu Val Leu Asp Leu Ser
            580                     585             590
Val Gln Gly Pro Gly Pro Val Leu Thr Val Leu Ala Leu Leu Leu Ala
            595                     600             605
Ser Thr Leu Ala
    610
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<210> 153
<211> 2111
<212> DNA
<213> Homo sapiens

<400> 153
ggccggccac tcccgtctgc tgtgacgcgc ggacagagag ctaccggtgg acccacggtg 60
cctccctccc tgggatctac acagaccatg gccttgccaa cggctcgacc cctgttgggg 120
tcctgtggga cccccgccct cggcagcctc ctgttcctgc tcttcagcct cggatgggtg 180
cagccctcga ggaccctggc tggagagaca gggcaggagg ctgcaccct ggacggagtc 240
ctggccaacc cacctaaacat ttccagcctc tccctcgcc aactccttgg cttcccgtgt 300
gcggaggtgt ccggcctgag cacggagcgt gtccgggagc tggctgtgc cttggcacag 360
aagaatgtca agctctcaac agagcagctg cgctgtctgg ctcaccggct ctctgagccc 420
cccgaggacc tggacgccct cccattggac ctgctgctat tcctcaaccc agatgcgttc 480
tcggggcccc aggcctgcac ccgtttcttc tcccgcatca cgaaggccaa tgtggacctg 540
ctcccgaggg gggctcccga gcgacagcgg ctgctgcctg cggctctggc ctgctggggt 600
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ccgggacccc tggaccagga ccagcaggag gcagccaggg cggctctgca gggcggggga 780
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cggttccggc gggaagtgga aagacagcc tgtccttcag gcaagaaggc ccgcgagata 1020
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ctaaagcata aactggatga gctctaccca caaggttacc ccgagtctgt gatccagcac 1200
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accctgaccg ccttctaccc tgggtacctg tgctccctca gccccgagga gctgagctcc 1440
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cagctggacg tcctctatcc caaggcccgc cttgctttcc agaacatgaa cgggtccgaa 1560
tacttcgtga gatccagtc cttcctgggt ggggcccca cggaggattt gaaggcgctc 1620
agtcagcaga atgtgagcat ggacttggcc acgttcatga agctgcggac ggatgcggtg 1680
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aagagaactc gcgctcagta aacgggaaca tgcccctgc agacacgtaa aaaaaaaaaa 2100
```

aaaaaaaaaa a   2111

<210> 154
<211> 622
<212> PRT
<213> Homo sapiens

<400> 154

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Met Ala Leu Pro Thr Ala Arg Pro Leu Leu Gly Ser Cys Gly Thr Pro
1               5                   10                  15
Ala Leu Gly Ser Leu Leu Phe Leu Leu Phe Ser Leu Gly Trp Val Gln
            20                  25                  30
Pro Ser Arg Thr Leu Ala Gly Glu Thr Gly Gln Glu Ala Ala Pro Leu
        35                  40                  45
Asp Gly Val Leu Ala Asn Pro Pro Asn Ile Ser Ser Leu Ser Pro Arg
    50                  55                  60
Gln Leu Leu Gly Phe Pro Cys Ala Glu Val Ser Gly Leu Ser Thr Glu
65                  70                  75                  80
Arg Val Arg Glu Leu Ala Val Ala Leu Ala Gln Lys Asn Val Lys Leu
                85                  90                  95
Ser Thr Glu Gln Leu Arg Cys Leu Ala His Arg Leu Ser Glu Pro Pro
            100                 105                 110
Glu Asp Leu Asp Ala Leu Pro Leu Asp Leu Leu Leu Phe Leu Asn Pro
        115                 120                 125
Asp Ala Phe Ser Gly Pro Gln Ala Cys Thr Arg Phe Phe Ser Arg Ile
    130                 135                 140
Thr Lys Ala Asn Val Asp Leu Leu Pro Arg Gly Ala Pro Glu Arg Gln
145                 150                 155                 160
Arg Leu Leu Pro Ala Ala Leu Ala Cys Trp Gly Val Arg Gly Ser Leu
                165                 170                 175
Leu Ser Glu Ala Asp Val Arg Ala Leu Gly Gly Leu Ala Cys Asp Leu
            180                 185                 190
Pro Gly Arg Phe Val Ala Glu Ser Ala Glu Val Leu Leu Pro Arg Leu
        195                 200                 205
Val Ser Cys Pro Gly Pro Leu Asp Gln Asp Gln Gln Glu Ala Ala Arg
    210                 215                 220
Ala Ala Leu Gln Gly Gly Gly Pro Pro Tyr Gly Pro Pro Ser Thr Trp
225                 230                 235                 240
Ser Val Ser Thr Met Asp Ala Leu Arg Gly Leu Leu Pro Val Leu Gly
                245                 250                 255
Gln Pro Ile Ile Arg Ser Ile Pro Gln Gly Ile Val Ala Ala Trp Arg
            260                 265                 270
Gln Arg Ser Ser Arg Asp Pro Ser Trp Arg Gln Pro Glu Arg Thr Ile
        275                 280                 285
Leu Arg Pro Arg Phe Arg Arg Glu Val Glu Lys Thr Ala Cys Pro Ser
    290                 295                 300
Gly Lys Lys Ala Arg Glu Ile Asp Glu Ser Leu Ile Phe Tyr Lys Lys
305                 310                 315                 320
Trp Glu Leu Glu Ala Cys Val Asp Ala Ala Leu Leu Ala Thr Gln Met
                325                 330                 335
Asp Arg Val Asn Ala Ile Pro Phe Thr Tyr Glu Gln Leu Asp Val Leu
            340                 345                 350
Lys His Lys Leu Asp Glu Leu Tyr Pro Gln Gly Tyr Pro Glu Ser Val
        355                 360                 365
Ile Gln His Leu Gly Tyr Leu Phe Leu Lys Met Ser Pro Glu Asp Ile
    370                 375                 380
Arg Lys Trp Asn Val Thr Ser Leu Glu Thr Leu Lys Ala Leu Leu Glu
385                 390                 395                 400
Val Asn Lys Gly His Glu Met Ser Pro Gln Val Ala Thr Leu Ile Asp
```

```
                    405                     410                     415
Arg Phe Val Lys Gly Arg Gly Gln Leu Asp Lys Asp Thr Leu Asp Thr
                420                     425                     430
Leu Thr Ala Phe Tyr Pro Gly Tyr Leu Cys Ser Leu Ser Pro Glu Glu
                435                     440                     445
Leu Ser Ser Val Pro Pro Ser Ser Ile Trp Ala Val Arg Pro Gln Asp
        450                     455                     460
Leu Asp Thr Cys Asp Pro Arg Gln Leu Asp Val Leu Tyr Pro Lys Ala
465                     470                     475                     480
Arg Leu Ala Phe Gln Asn Met Asn Gly Ser Glu Tyr Phe Val Lys Ile
                485                     490                     495
Gln Ser Phe Leu Gly Gly Ala Pro Thr Glu Asp Leu Lys Ala Leu Ser
                500                     505                     510
Gln Gln Asn Val Ser Met Asp Leu Ala Thr Phe Met Lys Leu Arg Thr
        515                     520                     525
Asp Ala Val Leu Pro Leu Thr Val Ala Glu Val Gln Lys Leu Leu Gly
        530                     535                     540
Pro His Val Glu Gly Leu Lys Ala Glu Glu Arg His Arg Pro Val Arg
545                     550                     555                     560
Asp Trp Ile Leu Arg Gln Arg Gln Asp Asp Leu Asp Thr Leu Gly Leu
                565                     570                     575
Gly Leu Gln Gly Gly Ile Pro Asn Gly Tyr Leu Val Leu Asp Leu Ser
                580                     585                     590
Val Gln Glu Ala Leu Ser Gly Thr Pro Cys Leu Leu Gly Pro Gly Pro
        595                     600                     605
Val Leu Thr Val Leu Ala Leu Leu Leu Ala Ser Thr Leu Ala
        610                     615                     620
```

<210> 155
<211> 1721
<212> DNA
<213> Homo sapiens

<400> 155
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acaccgggca cccagtctcc tttcttcctg ctgctgctcc tcacagtgct tacagttgtt 120
acaggttctg gtcatgcaag ctctacccca ggtggagaaa aggagacttc ggctacccag 180
agaagttcag tgcccagctc tactgagaag aatgctgtga gtatgaccag cagcgtactc 240
tccagccaca gccccggttc aggctcctcc accactcagg gacaggatgt cactctggcc 300
ccggccacgg aaccagcttc aggttcagct gccacctggg gacaggatgt cacctcggtc 360
ccagtcacca ggccagccct gggctccacc accccgccag cccacgatgt cacctcagcc 420
ccggacaaca agccagcccc gggctccacc gcccccccag cccacggtgt cacctcggcc 480
ccggacacca ggccgccccc gggctccacc gcccccccag cccacggtgt cacctcggcc 540
ccggacacca ggccgccccc gggctccacc gcgcccgcag cccacggtgt cacctcggcc 600
ccggacacca ggccggcccc gggctccacc gcccccccag cccatggtgt cacctcggcc 660
ccggacaaca ggcccgcctt ggcgtccacc gcccctccag tccacaatgt cacctcggcc 720
tcaggctctg catcaggctc agcttctact ctggtgcaca acggcacctc tgccagggct 780
accacaaccc cagccagcaa gagcactcca ttctcaattc ccagccacca ctctgatact 840
cctaccaccc ttgccagcca tagcaccaag actgatgcca gtagcactca ccatagcacg 900
gtacctcctc tcacctcctc caatcacagc acttctcccc agttgtctac tggggtctct 960
ttctttttcc tgtcttttca catttcaaac ctccagttta attcctctct ggaagatccc 1020
agcaccgact actaccaaga gctgcagaga gacatttctg aaatgttttt gcagatttat 1080
aaacaagggg gttttctggg cctctccaat attaagttca ggccaggatc tgtggtggta 1140
caattgactc tggccttccg agaaggtacc atcaatgtcc acgacgtgga gacacagttc 1200
aatcagtata aaacggaagc agcctctcga tataacctga cgatctcaga cgtcagcgtg 1260
agtgatgtgc catttccttt ctctgcccag tctggggctg gggtgccagg ctggggcatc 1320
gcgctgctgg tgctggtctg tgttctggtt gcgctggcca ttgtctatct cattgccttg 1380
gctgtctgtc agtgccgccg aaagaactac gggcagctgg acatctttcc agcccgggat 1440
```

```
acctaccatc ctatgagcga gtaccccacc taccacaccc atgggcgcta tgtgcccccт 1500
agcagtaccg atcgtagccc ctatgagaag gtttctgcag gtaatggtgg cagcagcctc 1560
tcttacacaa acccagcagt ggcagccact tctgccaact tgtaggggca cgtcgccctc 1620
tgagctgagt ggccagccag tgccattcca ctccactcag ggctctctgg gccagtcctc 1680
ctgggagccc ccaccacaac acttcccagg catggaattc c                     1721
```

<210> 156
<211> 515
<212> PRT
<213> Homo sapiens

<400> 156

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Met Thr Pro Gly Thr Gln Ser Pro Phe Phe Leu Leu Leu Leu Leu Thr
1               5                   10                  15
Val Leu Thr Val Val Thr Gly Ser Gly His Ala Ser Ser Thr Pro Gly
            20                  25                  30
Gly Glu Lys Glu Thr Ser Ala Thr Gln Arg Ser Ser Val Pro Ser Ser
        35                  40                  45
Thr Glu Lys Asn Ala Val Ser Met Thr Ser Ser Val Leu Ser Ser His
    50                  55                  60
Ser Pro Gly Ser Gly Ser Ser Thr Thr Gln Gly Gln Asp Val Thr Leu
65                  70                  75                  80
Ala Pro Ala Thr Glu Pro Ala Ser Gly Ser Ala Ala Thr Trp Gly Gln
            85                  90                  95
Asp Val Thr Ser Val Pro Val Thr Arg Pro Ala Leu Gly Ser Thr Thr
            100                 105                 110
Pro Pro Ala His Asp Val Thr Ser Ala Pro Asp Asn Lys Pro Ala Pro
        115                 120                 125
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    130                 135                 140
Arg Pro Pro Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
145                 150                 155                 160
Ala Pro Asp Thr Arg Pro Pro Pro Gly Ser Thr Ala Pro Ala Ala His
            165                 170                 175
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
            180                 185                 190
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Asn Arg Pro Ala Leu
        195                 200                 205
Ala Ser Thr Ala Pro Pro Val His Asn Val Thr Ser Ala Ser Gly Ser
    210                 215                 220
Ala Ser Gly Ser Ala Ser Thr Leu Val His Asn Gly Thr Ser Ala Arg
225                 230                 235                 240
Ala Thr Thr Thr Pro Ala Ser Lys Ser Thr Pro Phe Ser Ile Pro Ser
            245                 250                 255
His His Ser Asp Thr Pro Thr Thr Leu Ala Ser His Ser Thr Lys Thr
            260                 265                 270
Asp Ala Ser Ser Thr His His Ser Thr Val Pro Pro Leu Thr Ser Ser
        275                 280                 285
Asn His Ser Thr Ser Pro Gln Leu Ser Thr Gly Val Ser Phe Phe Phe
    290                 295                 300
Leu Ser Phe His Ile Ser Asn Leu Gln Phe Asn Ser Ser Leu Glu Asp
305                 310                 315                 320
Pro Ser Thr Asp Tyr Tyr Gln Glu Leu Gln Arg Asp Ile Ser Glu Met
            325                 330                 335
Phe Leu Gln Ile Tyr Lys Gln Gly Gly Phe Leu Gly Leu Ser Asn Ile
            340                 345                 350
Lys Phe Arg Pro Gly Ser Val Val Val Gln Leu Thr Leu Ala Phe Arg
        355                 360                 365
Glu Gly Thr Ile Asn Val His Asp Val Glu Thr Gln Phe Asn Gln Tyr
```

```
       370                         375                         380
Lys Thr Glu Ala Ala Ser Arg Tyr Asn Leu Thr Ile Ser Asp Val Ser
385                         390                         395                 400
Val Ser Asp Val Pro Phe Pro Phe Ser Ala Gln Ser Gly Ala Gly Val
                    405                         410                         415
Pro Gly Trp Gly Ile Ala Leu Leu Val Leu Val Cys Val Leu Val Ala
                    420                         425                         430
Leu Ala Ile Val Tyr Leu Ile Ala Leu Ala Val Cys Gln Cys Arg Arg
            435                         440                         445
Lys Asn Tyr Gly Gln Leu Asp Ile Phe Pro Ala Arg Asp Thr Tyr His
        450                         455                         460
Pro Met Ser Glu Tyr Pro Thr Tyr His Thr His Gly Arg Tyr Val Pro
465                         470                         475                 480
Pro Ser Ser Thr Asp Arg Ser Pro Tyr Glu Lys Val Ser Ala Gly Asn
                    485                         490                         495
Gly Gly Ser Ser Leu Ser Tyr Thr Asn Pro Ala Val Ala Ala Thr Ser
                    500                         505                         510
Ala Asn Leu
        515
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<210> 157
<211> 4139
<212> DNA
<213> Homo sapiens

<400> 157
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tttcaccacc accatgacac cgggcaccca gtctcctttc ttcctgctgc tgctcctcac 120
agtgcttaca gttgttacag gttctggtca tgcaagctct accccaggtg gagaaaagga 180
gacttcggct acccagagaa gttcagtgcc cagctctact gagaagaatg ctgtgagtat 240
gaccagcagc gtactctcca gccacagccc cggttcaggc tcctccacca ctcagggaca 300
ggatgtcact ctggccccgg ccacggaacc agcttcaggt tcagctgcca cctggggaca 360
ggatgtcacc tcggtcccag tcaccaggcc agccctgggc tccaccaccc cgccagccca 420
cgatgtcacc tcagccccgg acaacaagcc agccccgggc tccaccgccc cccagccca 480
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 540
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 600
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 660
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 720
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 780
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 840
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 900
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 960
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 1020
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cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 1260
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cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 1440
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cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 1560
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cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 1800
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 1860
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc cccagccca 1920
```

```
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc ccccagccca 1980
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc ccccagccca 2040
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc ccccagccca 2100
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc ccccagccca 2160
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc ccccagccca 2220
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc ccccagccca 2280
cggtgtcacc tcggccccgg acaccaggcc ggccccgggc tccaccgccc ccccagccca 2340
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cgtggagaca cagttcaatc agtataaaac ggaagcagcc tctcgatata acctgacgat 3480
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ctatctcatt gccttggctg tctgtcagtg ccgccgaaag aactacgggc agctggacat 3660
ctttccagcc cgggatacct accatcctat gagcgagtac cccacctacc acacccatgg 3720
gcgctatgtg ccccctagca gtaccgatcg tagcccctat gagaaggttt ctgcaggtaa 3780
cggtggcagc agcctctctt acacaaaccc agcagtggca gccgcttctg ccaacttgta 3840
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caggccagag ccctgcacc ctgtttgggctg tggtgagctg ggagttcagg tgggctgctc 3960
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gtgggcccct gaggctcatg cctgggaagt gttgtggggg ctcccaggag gactggccca 4080
gagagccctg agatagcggg gatcctgaac tggactgaat aaaacgtggt ctcccactg 4139
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<210> 158
<211> 1255
<212> PRT
<213> Homo sapiens

<400> 158
Met Thr Pro Gly Thr Gln Ser Pro Phe Phe Leu Leu Leu Leu Leu Thr
1               5                   10                  15
Val Leu Thr Val Val Thr Gly Ser Gly His Ala Ser Ser Thr Pro Gly
            20                  25                  30
Gly Glu Lys Glu Thr Ser Ala Thr Gln Arg Ser Ser Val Pro Ser Ser
        35                  40                  45
Thr Glu Lys Asn Ala Val Ser Met Thr Ser Ser Val Leu Ser Ser His
    50                  55                  60
Ser Pro Gly Ser Gly Ser Ser Thr Thr Gln Gly Gln Asp Val Thr Leu
65                  70                  75                  80
Ala Pro Ala Thr Glu Pro Ala Ser Gly Ser Ala Ala Thr Trp Gly Gln
                85                  90                  95
Asp Val Thr Ser Val Pro Val Thr Arg Pro Ala Leu Gly Ser Thr Thr
                100                 105                 110
Pro Pro Ala His Asp Val Thr Ser Ala Pro Asp Asn Lys Pro Ala Pro
```

```
            115                   120                   125
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    130                   135                   140
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
    145                   150                   155                   160
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                165                   170                   175
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
                180                   185                   190
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
                195                   200                   205
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    210                   215                   220
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
    225                   230                   235                   240
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                245                   250                   255
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
                260                   265                   270
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
                275                   280                   285
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    290                   295                   300
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
    305                   310                   315                   320
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                325                   330                   335
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
                340                   345                   350
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
                355                   360                   365
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    370                   375                   380
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
    385                   390                   395                   400
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                405                   410                   415
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
                420                   425                   430
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
                435                   440                   445
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    450                   455                   460
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
    465                   470                   475                   480
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                485                   490                   495
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
                500                   505                   510
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
                515                   520                   525
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    530                   535                   540
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
    545                   550                   555                   560
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                565                   570                   575
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
                580                   585                   590
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Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
        595   600   605
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    610   615   620
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
625   630   635   640
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                645   650   655
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
            660   665   670
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
        675   680   685
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    690   695   700
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
705   710   715   720
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                725   730   735
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
            740   745   750
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
        755   760   765
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    770   775   780
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
785   790   795   800
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                805   810   815
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
            820   825   830
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
        835   840   845
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr
    850   855   860
Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser
865   870   875   880
Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala Pro Pro Ala His
                885   890   895
Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro Gly Ser Thr Ala
            900   905   910
Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Thr Arg Pro Ala Pro
        915   920   925
Gly Ser Thr Ala Pro Pro Ala His Gly Val Thr Ser Ala Pro Asp Asn
    930   935   940
Arg Pro Ala Leu Gly Ser Thr Ala Pro Pro Val His Asn Val Thr Ser
945   950   955   960
Ala Ser Gly Ser Ala Ser Gly Ser Ala Ser Thr Leu Val His Asn Gly
                965   970   975
Thr Ser Ala Arg Ala Thr Thr Pro Ala Ser Lys Ser Thr Pro Phe
            980   985   990
Ser Ile Pro Ser His His Ser Asp Thr Pro Thr Thr Leu Ala Ser His
        995   1000   1005
Ser Thr Lys Thr Asp Ala Ser Ser Thr His His Ser Ser Val Pro Pro
    1010   1015   1020
Leu Thr Ser Ser Asn His Ser Thr Ser Pro Gln Leu Ser Thr Gly Val
1025   1030   1035   1040
Ser Phe Phe Phe Leu Ser Phe His Ile Ser Asn Leu Gln Phe Asn Ser
            1045   1050   1055
Ser Leu Glu Asp Pro Ser Thr Asp Tyr Tyr Gln Glu Leu Gln Arg Asp

```
          1060                   1065                   1070
Ile Ser Glu Met Phe Leu Gln Ile Tyr Lys Gln Gly Gly Phe Leu Gly
          1075                   1080                   1085
Leu Ser Asn Ile Lys Phe Arg Pro Gly Ser Val Val Val Gln Leu Thr
     1090                   1095                   1100
Leu Ala Phe Arg Glu Gly Thr Ile Asn Val His Asp Val Glu Thr Gln
1105                   1110                   1115                   1120
Phe Asn Gln Tyr Lys Thr Glu Ala Ala Ser Arg Tyr Asn Leu Thr Ile
               1125                   1130                   1135
Ser Asp Val Ser Val Ser Asp Val Pro Phe Pro Phe Ser Ala Gln Ser
          1140                   1145                   1150
Gly Ala Gly Val Pro Gly Trp Gly Ile Ala Leu Leu Val Leu Val Cys
          1155                   1160                   1165
Val Leu Val Ala Leu Ala Ile Val Tyr Leu Ile Ala Leu Ala Val Cys
     1170                   1175                   1180
Gln Cys Arg Arg Lys Asn Tyr Gly Gln Leu Asp Ile Phe Pro Ala Arg
1185                   1190                   1195                   1200
Asp Thr Tyr His Pro Met Ser Glu Tyr Pro Thr Tyr His Thr His Gly
               1205                   1210                   1215
Arg Tyr Val Pro Pro Ser Ser Thr Asp Arg Ser Pro Tyr Glu Lys Val
          1220                   1225                   1230
Ser Ala Gly Asn Gly Gly Ser Ser Leu Ser Tyr Thr Asn Pro Ala Val
          1235                   1240                   1245
Ala Ala Ala Ser Ala Asn Leu
     1250                   1255
```

<210> 159
<211> 2627
<212> DNA
<213> Homo sapiens

<400> 159

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ccggcccggc tcccgctccg ggctctgccg gcgggcgggc gagcgcggcg cggtccgggc 120
cggggggatg tctcggcgga cgcgctgcga ggatctggat gagctgcact accaggacac 180
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tccagacctt gtcaaggggc catggaccaa agaggaagac caaaaagtca tcgagctggt 420
taagaagtat ggcacaaagc agtggacact gattgccaag cacctgaagg ccggctggg 480
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gaccgaggag gaggaccgca tcatctgcga ggcccacaag gtgctgggca accgctgggc 600
cgagatcgcc aagatgttgc agggaggac agacaatgct gtgaagaatc actggaactc 660
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cccagtgtac ttgctgctgg agctcgagga caaggacggc ctccagagtg cccagcccac 780
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224

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<213> Homo sapiens

<400> 160

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Thr Asp Gln Gln Cys Gln Tyr Arg Trp Leu Arg Val Leu Asn Pro Asp
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Ala Lys Met Leu Pro Gly Arg Thr Asp Asn Ala Val Lys Asn His Trp
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Asn Ser Thr Ile Lys Arg Lys Val Asp Thr Gly Gly Phe Leu Ser Glu
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Ser Lys Asp Cys Lys Pro Pro Val Tyr Leu Leu Leu Glu Leu Glu Asp
            195                 200                 205
Lys Asp Gly Leu Gln Ser Ala Gln Pro Thr Glu Gly Gln Gly Ser Leu
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Leu Thr Asn Trp Pro Ser Val Pro Pro Thr Ile Lys Glu Glu Glu Asn
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Ser Glu Glu Glu Leu Ala Ala Ala Thr Thr Ser Lys Glu Gln Glu Pro
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Ile Gly Thr Asp Leu Asp Ala Val Arg Thr Pro Glu Pro Leu Glu Glu
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Glu Asp Ser Ile Asn Asn Ser Leu Val Gln Leu Gln Ala Ser His Gln
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385   390   395   400
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Leu Asp Ser Cys Asn Ser Leu Thr Pro Lys Ser Thr Pro Val Lys Thr
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Pro Thr Thr Ala Pro Ser Asn Ser Ser Ser Leu Thr Leu Ser Gly Ile
        610   615   620
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Glu Lys Ala Ala Val Ala Gln Lys Pro Arg Ser His Phe Thr Thr Pro
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Ala Pro Met Ser Ser Ala Trp Lys Thr Val Ala Cys Gly Gly Thr Arg
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<213> Homo sapiens

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<212> PRT
<213> Homo sapiens

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Lys Arg Leu Val Trp Val Pro Ser Glu Lys Gln Gly Phe Glu Ala Ala
            35                  40                  45
Ser Ile Lys Glu Glu Lys Gly Asp Glu Val Val Val Glu Leu Val Glu
    50                  55                  60
Asn Gly Lys Lys Val Thr Val Gly Lys Asp Asp Ile Gln Lys Met Asn
65                  70                  75                  80
Pro Pro Lys Phe Ser Lys Val Glu Asp Met Ala Glu Leu Thr Cys Leu
                85                  90                  95
Asn Glu Ala Ser Val Leu His Asn Leu Arg Glu Arg Tyr Phe Ser Gly
            100                 105                 110
Leu Ile Tyr Thr Tyr Ser Gly Leu Phe Cys Val Val Val Asn Pro Tyr
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Lys His Leu Pro Ile Tyr Ser Glu Lys Ile Val Asp Met Tyr Lys Gly
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Lys Lys Arg His Glu Met Pro Pro His Ile Tyr Ala Ile Ala Asp Thr
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Ala Tyr Arg Ser Met Leu Gln Asp Arg Glu Asp Gln Ser Ile Leu Cys
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Thr Gly Glu Ser Gly Ala Gly Lys Thr Glu Asn Thr Lys Lys Val Ile
            180                 185                 190
Gln Tyr Leu Ala Val Val Ala Ser Ser His Lys Gly Lys Lys Asp Thr
            195                 200                 205
Ser Ile Thr Gly Glu Leu Glu Lys Gln Leu Leu Gln Ala Asn Pro Ile
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225                 230                 235                 240
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    370                 375                 380
Met Gly Ile Asn Val Thr Asp Phe Thr Arg Ser Ile Leu Thr Pro Arg
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229

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Ile Phe Glu Val Asn Ser Phe Glu Gln Leu Cys Ile Asn Tyr Thr Asn
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Lys Glu Arg Gln Gln Lys Ala Glu Asn Glu Leu Lys Glu Leu Glu Gln
865             870             875             880
Lys His Ser Gln Leu Thr Glu Glu Lys Asn Leu Leu Gln Glu Gln Leu
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Ala Arg Leu Glu Glu Glu Glu Asp Arg Gly Gln Gln Leu Gln Ala Glu
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Arg Lys Lys Met Ala Gln Gln Met Leu Asp Leu Glu Glu Gln Leu Glu
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Glu Glu Glu Ala Ala Arg Gln Lys Leu Gln Leu Glu Lys Val Thr Ala
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Glu Ala Lys Ile Lys Lys Leu Glu Asp Glu Ile Leu Val Met Asp Asp
               980                 985                 990
Gln Asn Asn Lys Leu Ser Lys Glu Arg Lys Leu Leu Glu Glu Arg Ile
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Ser Asp Leu Thr Thr Asn Leu Ala Glu Glu Glu Glu Lys Ala Lys Asn
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Leu Thr Lys Leu Lys Asn Lys His Glu Ser Met Ile Ser Glu Leu Glu
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Val Arg Leu Lys Lys Glu Glu Lys Ser Arg Gln Glu Leu Glu Lys Leu
               1045                1050                1055
Lys Arg Lys Leu Glu Gly Asp Ala Ser Asp Phe His Glu Gln Ile Ala
               1060                1065                1070
Asp Leu Gln Ala Gln Ile Ala Glu Leu Lys Met Gln Leu Ala Lys Lys
           1075                1080                1085
Glu Glu Glu Leu Gln Ala Ala Leu Ala Arg Leu Asp Asp Glu Ile Ala
       1090                1095                1100
Gln Lys Asn Asn Ala Leu Lys Lys Ile Arg Glu Leu Glu Gly His Ile
1105                1110                1115                1120
Ser Asp Leu Gln Glu Asp Leu Asp Ser Glu Arg Ala Ala Arg Asn Lys
               1125                1130                1135
Ala Glu Lys Gln Lys Arg Asp Leu Gly Glu Glu Leu Glu Ala Leu Lys
           1140                1145                1150
Thr Glu Leu Glu Asp Thr Leu Asp Ser Thr Ala Thr Gln Gln Glu Leu
       1155                1160                1165
Arg Ala Lys Arg Glu Gln Glu Val Thr Val Leu Lys Lys Ala Leu Asp
       1170                1175                1180
Glu Glu Thr Arg Ser His Glu Ala Gln Val Gln Glu Met Arg Gln Lys
1185                1190                1195                1200
His Ala Gln Ala Val Glu Glu Leu Thr Glu Gln Leu Glu Gln Phe Lys
           1205                1210                1215
Arg Ala Lys Ala Asn Leu Asp Lys Asn Lys Gln Thr Leu Glu Lys Glu
           1220                1225                1230
Asn Ala Asp Leu Ala Gly Glu Leu Arg Val Leu Gly Gln Ala Lys Gln
           1235                1240                1245
Glu Val Glu His Lys Lys Lys Lys Leu Glu Ala Gln Val Gln Glu Leu
       1250                1255                1260
Gln Ser Lys Cys Ser Asp Gly Glu Arg Ala Arg Ala Glu Leu Asn Asp
1265                1270                1275                1280
Lys Val His Lys Leu Gln Asn Glu Val Glu Ser Val Thr Gly Met Leu
               1285                1290                1295
Asn Glu Ala Glu Gly Lys Ala Ile Lys Leu Ala Lys Asp Val Ala Ser
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Leu Ser Ser Gln Leu Gln Asp Thr Gln Glu Leu Leu Gln Glu Glu Thr
       1315                1320                1325
Arg Gln Lys Leu Asn Val Ser Thr Lys Leu Arg Gln Leu Glu Glu Glu
       1330                1335                1340
Arg Asn Ser Leu Gln Asp Gln Leu Asp Glu Glu Met Glu Ala Lys Gln
1345                1350                1355                1360
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Gly Lys Lys Arg Phe Gln Lys Glu Ile Glu Asn Leu Thr Gln Gln Tyr
            1395                1400                1405
Glu Glu Lys Ala Ala Ala Tyr Asp Lys Leu Glu Lys Thr Lys Asn Arg
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Leu Gln Gln Glu Leu Asp Asp Leu Val Val Asp Leu Asp Asn Gln Arg
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Gln Leu Val Ser Asn Leu Glu Lys Lys Gln Arg Lys Phe Asp Gln Leu
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            1460                1465                1470
Arg Ala Glu Ala Glu Ala Arg Glu Lys Glu Thr Lys Ala Leu Ser Leu
            1475                1480                1485
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Thr Asn Lys Met Leu Lys Ala Glu Met Glu Asp Leu Val Ser Ser Lys
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Asp Asp Val Gly Lys Asn Val His Glu Leu Glu Lys Ser Lys Arg Ala
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Leu Glu Thr Gln Met Glu Glu Met Lys Thr Gln Leu Glu Glu Leu Glu
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        1570                1575                1580
Glu Gln Asn Glu Glu Lys Arg Arg Gln Leu Gln Arg Gln Leu His Glu
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Tyr Glu Thr Glu Leu Glu Asp Glu Arg Lys Gln Arg Ala Leu Ala Ala
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Ala Ala Lys Lys Lys Leu Glu Gly Asp Leu Lys Asp Leu Glu Leu Gln
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Ala Asp Ser Ala Ile Lys Gly Arg Glu Glu Ala Ile Lys Gln Leu Arg
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Lys Leu Gln Ala Gln Met Lys Asp Phe Gln Arg Glu Leu Glu Asp Ala
    1650                1655                1660
Arg Ala Ser Arg Asp Glu Ile Phe Ala Thr Ala Lys Glu Asn Glu Lys
1665                1670                1675                1680
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            1685                1690                1695
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Leu Ala Glu Glu Leu Ala Ser Ser Leu Ser Gly Arg Asn Ala Leu Gln
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Asp Glu Lys Arg Arg Leu Glu Ala Arg Ile Ala Gln Leu Glu Glu Glu
    1730                1735                1740
Leu Glu Glu Glu Gln Gly Asn Met Glu Ala Met Ser Asp Arg Val Arg
1745                1750                1755                1760
Lys Ala Thr Gln Gln Ala Glu Gln Leu Ser Asn Glu Leu Ala Thr Glu
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Arg Ser Thr Ala Gln Lys Asn Glu Ser Ala Arg Gln Gln Leu Glu Arg
            1780                1785                1790
Gln Asn Lys Glu Leu Arg Ser Lys Leu His Glu Met Glu Gly Ala Val
        1795                1800                1805
Lys Ser Lys Phe Lys Ser Thr Ile Ala Ala Leu Glu Ala Lys Ile Ala
    1810                1815                1820
Gln Leu Glu Glu Gln Val Glu Gln Glu Ala Arg Glu Lys Gln Ala Ala
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1825                     1830                1835                     1840
Thr Lys Ser Leu Lys Gln Lys Asp Lys Lys Leu Lys Glu Ile Leu Leu
                1845                1850                     1855
Gln Val Glu Asp Glu Arg Lys Met Ala Glu Gln Tyr Lys Glu Gln Ala
            1860                1865                     1870
Glu Lys Gly Asn Ala Arg Val Lys Gln Leu Lys Arg Gln Leu Glu Glu
        1875                1880                     1885
Ala Glu Glu Glu Ser Gln Arg Ile Asn Ala Asn Arg Arg Lys Leu Gln
    1890                1895                     1900
Arg Glu Leu Asp Glu Ala Thr Glu Ser Asn Glu Ala Met Gly Arg Glu
1905                    1910                1915                     1920
Val Asn Ala Leu Lys Ser Lys Leu Arg Arg Gly Asn Glu Thr Ser Phe
                1925                1930                     1935
Val Pro Ser Arg Arg Ser Gly Gly Arg Arg Val Ile Glu Asn Ala Asp
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        1955                1960                     1965
Lys Ala Ser Glu
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<211> 6900
<212> DNA
<213> Homo sapiens

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<211> 1938
<212> PRT
<213> Homo sapiens

<400> 164
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            20                  25                  30
Lys Arg Leu Val Trp Val Pro Ser Glu Lys Gln Gly Phe Glu Ala Ala
        35                  40                  45
Ser Ile Lys Glu Glu Lys Gly Asp Glu Val Val Val Glu Leu Val Glu
    50                  55                  60
Asn Gly Lys Lys Val Thr Val Gly Lys Asp Asp Ile Gln Lys Met Asn
65                  70                  75                  80
Pro Pro Lys Phe Ser Lys Val Glu Asp Met Ala Glu Leu Thr Cys Leu
                85                  90                  95
Asn Glu Ala Ser Val Leu His Asn Leu Arg Glu Arg Tyr Phe Ser Gly
            100                 105                 110
Leu Ile Tyr Thr Tyr Ser Gly Leu Phe Cys Val Val Val Asn Pro Tyr
        115                 120                 125
Lys His Leu Pro Ile Tyr Ser Glu Lys Ile Val Asp Met Tyr Lys Gly
    130                 135                 140
Lys Lys Arg His Glu Met Pro Pro His Ile Tyr Ala Ile Ala Asp Thr
145                 150                 155                 160
Ala Tyr Arg Ser Met Leu Gln Asp Arg Glu Asp Gln Ser Ile Leu Cys
                165                 170                 175
Thr Gly Glu Ser Gly Ala Gly Lys Thr Glu Asn Thr Lys Lys Val Ile
            180                 185                 190
Gln Tyr Leu Ala Val Val Ala Ser Ser His Lys Gly Lys Lys Asp Thr
        195                 200                 205
Ser Ile Thr Gly Glu Leu Glu Lys Gln Leu Leu Gln Ala Asn Pro Ile
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Leu Glu Ala Phe Gly Asn Ala Lys Thr Val Lys Asn Asp Asn Ser Ser
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                        245                     250                     255
Val Gly Ala Asn Ile Glu Thr Tyr Leu Leu Glu Lys Ser Arg Ala Ile
                260                     265                     270
Arg Gln Ala Arg Asp Glu Arg Thr Phe His Ile Phe Tyr Tyr Met Ile
            275                     280                     285
Ala Gly Ala Lys Glu Lys Met Arg Ser Asp Leu Leu Leu Glu Gly Phe
            290                     295                     300
Asn Asn Tyr Thr Phe Leu Ser Asn Gly Phe Val Pro Ile Pro Ala Ala
305                     310                     315                     320
Gln Asp Asp Glu Met Phe Gln Glu Thr Val Glu Ala Met Ala Ile Met
                325                     330                     335
Gly Phe Ser Glu Glu Glu Gln Leu Ser Ile Leu Lys Val Val Ser Ser
                340                     345                     350
Val Leu Gln Leu Gly Asn Ile Val Phe Lys Lys Glu Arg Asn Thr Asp
            355                     360                     365
Gln Ala Ser Met Pro Asp Asn Thr Ala Ala Gln Lys Val Cys His Leu
            370                     375                     380
Met Gly Ile Asn Val Thr Asp Phe Thr Arg Ser Ile Leu Thr Pro Arg
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Ile Lys Val Gly Arg Asp Val Val Gln Lys Ala Gln Thr Lys Glu Gln
                405                     410                     415
Ala Asp Phe Ala Val Glu Ala Leu Ala Lys Ala Thr Tyr Glu Arg Leu
            420                     425                     430
Phe Arg Trp Ile Leu Thr Arg Val Asn Lys Ala Leu Asp Lys Thr His
            435                     440                     445
Arg Gln Gly Ala Ser Phe Leu Gly Ile Leu Asp Ile Ala Gly Phe Glu
            450                     455                     460
Ile Phe Glu Val Asn Ser Phe Glu Gln Leu Cys Ile Asn Tyr Thr Asn
465                     470                     475                     480
Glu Lys Leu Gln Gln Leu Phe Asn His Thr Met Phe Ile Leu Glu Gln
                485                     490                     495
Glu Glu Tyr Gln Arg Glu Gly Ile Glu Trp Asn Phe Ile Asp Phe Gly
            500                     505                     510
Leu Asp Leu Gln Pro Cys Ile Glu Leu Ile Glu Arg Pro Asn Asn Pro
            515                     520                     525
Pro Gly Val Leu Ala Leu Leu Asp Glu Glu Cys Trp Phe Pro Lys Ala
            530                     535                     540
Thr Asp Lys Ser Phe Val Glu Lys Leu Cys Thr Glu Gln Gly Ser His
545                     550                     555                     560
Pro Lys Phe Gln Lys Pro Lys Gln Leu Lys Asp Lys Thr Glu Phe Ser
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Ile Ile His Tyr Ala Gly Lys Val Asp Tyr Asn Ala Ser Ala Trp Leu
                580                     585                     590
Thr Lys Asn Met Asp Pro Leu Asn Asp Asn Val Thr Ser Leu Leu Asn
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Ala Ser Ser Asp Lys Phe Val Ala Asp Leu Trp Lys Asp Val Asp Arg
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Ile Val Gly Leu Asp Gln Met Ala Lys Met Thr Glu Ser Ser Leu Pro
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Ser Ala Ser Lys Thr Lys Lys Gly Met Phe Arg Thr Val Gly Gln Leu
                645                     650                     655
Tyr Lys Glu Gln Leu Gly Lys Leu Met Thr Thr Leu Arg Asn Thr Thr
                660                     665                     670
Pro Asn Phe Val Arg Cys Ile Ile Pro Asn His Glu Lys Arg Ser Gly
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Leu Glu Gly Ile Arg Ile Cys Arg Gln Gly Phe Pro Asn Arg Ile Val
705             710             715             720
Phe Gln Glu Phe Arg Gln Arg Tyr Glu Ile Leu Ala Ala Asn Ala Ile
            725             730             735
Pro Lys Gly Phe Met Asp Gly Lys Gln Ala Cys Ile Leu Met Ile Lys
            740             745             750
Ala Leu Glu Leu Asp Pro Asn Leu Tyr Arg Ile Gly Gln Ser Lys Ile
            755             760             765
Phe Phe Arg Thr Gly Val Leu Ala His Leu Glu Glu Glu Arg Asp Leu
    770             775             780
Lys Ile Thr Asp Val Ile Met Ala Phe Gln Ala Met Cys Arg Gly Tyr
785             790             795             800
Leu Ala Arg Lys Ala Phe Ala Lys Arg Gln Gln Gln Leu Thr Ala Met
            805             810             815
Lys Val Ile Gln Arg Asn Cys Ala Ala Tyr Leu Lys Leu Arg Asn Trp
            820             825             830
Gln Trp Trp Arg Leu Phe Thr Lys Val Lys Pro Leu Leu Gln Val Thr
            835             840             845
Arg Gln Glu Glu Glu Met Gln Ala Lys Glu Asp Glu Leu Gln Lys Thr
    850             855             860
Lys Glu Arg Gln Gln Lys Ala Glu Asn Glu Leu Lys Glu Leu Glu Gln
865             870             875             880
Lys His Ser Gln Leu Thr Glu Glu Lys Asn Leu Leu Gln Glu Gln Leu
            885             890             895
Gln Ala Glu Thr Glu Leu Tyr Ala Glu Ala Glu Glu Met Arg Val Arg
            900             905             910
Leu Ala Ala Lys Lys Gln Glu Leu Glu Glu Ile Leu His Glu Met Glu
            915             920             925
Ala Arg Leu Glu Glu Glu Glu Asp Arg Gly Gln Gln Leu Gln Ala Glu
    930             935             940
Arg Lys Lys Met Ala Gln Gln Met Leu Asp Leu Glu Glu Gln Leu Glu
945             950             955             960
Glu Glu Glu Ala Ala Arg Gln Lys Leu Gln Leu Glu Lys Val Thr Ala
            965             970             975
Glu Ala Lys Ile Lys Lys Leu Glu Asp Glu Ile Leu Val Met Asp Asp
            980             985             990
Gln Asn Asn Lys Leu Ser Lys Glu Arg Lys Leu Leu Glu Glu Arg Ile
    995             1000            1005
Ser Asp Leu Thr Thr Asn Leu Ala Glu Glu Glu Lys Ala Lys Asn
    1010            1015            1020
Leu Thr Lys Leu Lys Asn Lys His Glu Ser Met Ile Ser Glu Leu Glu
1025            1030            1035            1040
Val Arg Leu Lys Lys Glu Glu Lys Ser Arg Gln Glu Leu Glu Lys Leu
            1045            1050            1055
Lys Arg Lys Leu Glu Gly Asp Ala Ser Asp Phe His Glu Gln Ile Ala
            1060            1065            1070
Asp Leu Gln Ala Gln Ile Ala Glu Leu Lys Met Gln Leu Ala Lys Lys
            1075            1080            1085
Glu Glu Glu Leu Gln Ala Ala Leu Ala Arg Leu Asp Asp Glu Ile Ala
    1090            1095            1100
Gln Lys Asn Asn Ala Leu Lys Lys Ile Arg Glu Leu Glu Gly His Ile
1105            1110            1115            1120
Ser Asp Leu Gln Glu Asp Leu Asp Ser Glu Arg Ala Ala Arg Asn Lys
            1125            1130            1135
Ala Glu Lys Gln Lys Arg Asp Leu Gly Glu Glu Leu Glu Ala Leu Lys
            1140            1145            1150
Thr Glu Leu Glu Asp Thr Leu Asp Ser Thr Ala Thr Gln Gln Glu Leu
```

```
          1155                    1160                    1165
Arg Ala Lys Arg Glu Gln Glu Val Thr Val Leu Lys Lys Ala Leu Asp
    1170                    1175                  '1180
Glu Glu Thr Arg Ser His Glu Ala Gln Val Gln Glu Met Arg Gln Lys
1185                    1190                    1195                    1200
His Ala Gln Ala Val Glu Glu Leu Thr Glu Gln Leu Glu Gln Phe Lys
                    1205                    1210                    1215·
Arg Ala Lys Ala Asn Leu Asp Lys Asn Lys Gln Thr Leu Glu Lys Glu
                    1220                    1225                    1230
Asn Ala Asp Leu Ala Gly Glu Leu Arg Val Leu Gly Gln Ala Lys Gln
                    1235                    1240                    1245
Glu Val Glu His Lys Lys Lys Lys Leu Glu Ala Gln Val Gln Glu Leu
                    1250                    1255                    1260
Gln Ser Lys Cys Ser Asp Gly Glu Arg Ala Arg Ala Glu Leu Asn Asp
1265                    1270                    1275                    1280
Lys Val His Lys Leu Gln Asn Glu Val Glu Ser Val Thr Gly Met Leu
                    1285                    1290                    1295
Asn Glu Ala Glu Gly Lys Ala Ile Lys Leu Ala Lys Asp Val Ala Ser
                    1300                    1305                    1310
Leu Ser Ser Gln Leu Gln Asp Thr Gln Glu Leu Leu Gln Glu Glu Thr
                    1315                    1320                    1325
Arg Gln Lys Leu Asn Val Ser Thr Lys Leu Arg Gln Leu Glu Glu Glu
                    1330                    1335                    1340
Arg Asn Ser Leu Gln Asp Gln Leu Asp Glu Glu Met Glu Ala Lys Gln
1345                    1350                    1355                    1360
Asn Leu Glu Arg His Ile Ser Thr Leu Asn Ile Gln Leu Ser Asp Ser
                    1365                    1370                    1375
Lys Lys Lys Leu Gln Asp Phe Ala Ser Thr Val Glu Ala Leu Glu Glu
                    1380                    1385                    1390
Gly Lys Lys Arg Phe Gln Lys Glu Ile Glu Asn Leu Thr Gln Gln Tyr
                    1395                    1400                    1405
Glu Glu Lys Ala Ala Ala Tyr Asp Lys Leu Glu Lys Thr Lys Asn Arg
    1410 ·                   1415                    1420
Leu Gln Gln Glu Leu Asp Asp Leu Val Val Asp Leu Asp Asn Gln Arg
1425                    1430                    1435                    1440
Gln Leu Val Ser Asn Leu Glu Lys Lys Gln Arg Lys Phe Asp Gln Leu
                    1445                    1450                    1455
Leu Ala Glu Glu Lys Asn Ile Ser Ser Lys Tyr Ala Asp Glu Arg Asp
                    1460                    1465                    1470
Arg Ala Glu Ala Glu Ala Arg Glu Lys Glu Thr Lys Ala Leu Ser Leu
                    1475                    1480                    1485
Ala Arg Ala Leu Glu Glu Ala Leu Glu Ala Lys Glu Glu Leu Glu Arg
    1490                    1495                    1500
Thr Asn Lys Met Leu Lys Ala Glu Met Glu Asp Leu Val Ser Ser Lys
1505                    1510                    1515                    1520
Asp Asp Val Gly Lys Asn Val His Glu Leu Glu Lys Ser Lys Arg Ala
                    1525                    1530                    1535
Leu Glu Thr Gln Met Glu Glu Met Lys Thr Gln Leu Glu Glu Leu Glu
                    1540                    1545                    1550
Asp Glu Leu Gln Ala Thr Glu Asp Ala Lys Leu Arg Leu Glu Val Asn
                    1555                    1560                    1565
Met Gln Ala Leu Lys Gly Gln Phe Glu Arg Asp Leu Gln Ala Arg Asp
    1570                    1575                    1580
Glu Gln Asn Glu Glu Lys Arg Arg Gln Leu Gln Arg Gln Leu His Glu
1585                    1590                    1595                    1600
Tyr Glu Thr Glu Leu Glu Asp Glu Arg Lys Gln Arg Ala Leu Ala Ala
                    1605                    1610                    1615
Ala Ala Lys Lys Lys Leu Glu Gly Asp Leu Lys Asp Leu Glu Leu Gln
                    1620                    1625                    1630
```

```
Ala Asp Ser Ala Ile Lys Gly Arg Glu Glu Ala Ile Lys Gln Leu Arg
        1635                1640                1645
Lys Leu Gln Ala Gln Met Lys Asp Phe Gln Arg Glu Leu Glu Asp Ala
        1650                1655                1660
Arg Ala Ser Arg Asp Glu Ile Phe Ala Thr Ala Lys Glu Asn Glu Lys
1665                1670                1675                1680
Lys Ala Lys Ser Leu Glu Ala Asp Leu Met Gln Leu Gln Glu Asp Leu
                1685                1690                1695
Ala Ala Ala Glu Arg Ala Arg Lys Gln Ala Asp Leu Glu Lys Glu Glu
                1700                1705                1710
Leu Ala Glu Glu Leu Ala Ser Ser Leu Ser Gly Arg Asn Ala Leu Gln
                1715                1720                1725
Asp Glu Lys Arg Arg Leu Glu Ala Arg Ile Ala Gln Leu Glu Glu Glu
        1730                1735                1740
Leu Glu Glu Glu Gln Gly Asn Met Glu Ala Met Ser Asp Arg Val Arg
1745                1750                1755                1760
Lys Ala Thr Gln Gln Ala Glu Gln Leu Ser Asn Glu Leu Ala Thr Glu
                1765                1770                1775
Arg Ser Thr Ala Gln Lys Asn Glu Ser Ala Arg Gln Gln Leu Glu Arg
        1780                1785                1790
Gln Asn Lys Glu Leu Arg Ser Lys Leu His Glu Met Glu Gly Ala Val
        1795                . 1800                1805
Lys Ser Lys Phe Lys Ser Thr Ile Ala Ala Leu Glu Ala Lys Ile Ala
        1810                1815                1820
Gln Leu Glu Glu Gln Val Glu Gln Glu Ala Arg Glu Lys Gln Ala Ala
1825                1830                1835                1840
Thr Lys Ser Leu Lys Gln Lys Asp Lys Lys Leu Lys Glu Ile Leu Leu
                1845                1850                1855
Gln Val Glu Asp Glu Arg Lys Met Ala Glu Gln Tyr Lys Glu Gln Ala
                1860                1865                1870
Glu Lys Gly Asn Ala Arg Val Lys Gln Leu Lys Arg Gln Leu Glu Glu
        1875                1880                1885
Ala Glu Glu Glu Ser Gln Arg Ile Asn Ala Asn Arg Arg Lys Leu Gln
        1890                1895                1900
Arg Glu Leu Asp Glu Ala Thr Glu Ser Asn Glu Ala Met Gly Arg Glu
1905                1910                1915                1920
Val Asn Ala Leu Lys Ser Lys Leu Arg Gly Pro Pro Pro Gln Glu Thr
                1925                1930                1935
Ser Gln
```

```
<210> 165
<211> 958
<212> DNA
<213> Homo sapiens

<400> 165
tctaaagctc agtggagctg ggtcatctca ggccttggct ccttgaactt ttggccgcca  60
tgtgcttccc gaaggtcctc tctgatgaca tgaagaagct gaaggcccga atggtaatgc 120
tcctccctac ttctgctcag gggttggggg cctgggtctc agcgtgtgac actgaggaca 180
ctgtgggaca cctgggaccc tggagggaca aggatccggc cctttggtgc caactctgcc 240
tctcttcaca gcaccaggcc atagaaagat tttatgataa aatgcaaaat gcagaatcag 300
gacgtggaca ggtgatgtcg agcctggcag agctggagga cgacttcaaa gagggctacc 360
tggagacagt ggcggcttat tatgaggagc agcacccaga gctcactcct ctacttgaaa 420
aagaaagaga tggattacgg tgccgaggca acagatcccc tgtcccggat gttgaggatc 480
ccgcaaccga ggagcctggg gagagctttt gtracaaggt catgagatgg ttccaggcca 540
tgctgcagcg gctgcagacc tggtggcacg gggttctggc ctgggtgaag gagaaggtgg 600
tggccctggt ccatgcagtg caggccctct ggaaacagtt ccagagtttc tgctgctctc 660
```

```
tgtcagagct cttcatgtcc tctttccagt cctacggagc cccacggggg gacaaggagg 720
agctgacacc ccagaagtgc tctgaacccc aatcctcaaa atgaagatac tgacaccacc 780
tttgccctcc ccgtcaccgc gcacccaccc tgacccctcc ctcagctgtc ctgtgccccg 840
ccctctcccg cacactcagt ccccctgcct ggcgttcctg ccgcagctct gacctggtgc 900
tgtcgccctg gcatcttaat aaaacctgct tatacttccc tggcagggag ataccatg   958
```

<210> 166
<211> 234
<212> PRT
<213> Homo sapiens

<400> 166

```
Met Cys Phe Pro Lys Val Leu Ser Asp Asp Met Lys Lys Leu Lys Ala
 1               5                  10                  15
Arg Met Val Met Leu Leu Pro Thr Ser Ala Gln Gly Leu Gly Ala Trp
            20                  25                  30
Val Ser Ala Cys Asp Thr Glu Asp Thr Val Gly His Leu Gly Pro Trp
            35                  40                  45
Arg Asp Lys Asp Pro Ala Leu Trp Cys Gln Leu Cys Leu Ser Ser Gln
        50                  55                  60
His Gln Ala Ile Glu Arg Phe Tyr Asp Lys Met Gln Asn Ala Glu Ser
65                  70                  75                  80
Gly Arg Gly Gln Val Met Ser Ser Leu Ala Glu Leu Glu Asp Asp Phe
                85                  90                  95
Lys Glu Gly Tyr Leu Glu Thr Val Ala Ala Tyr Tyr Glu Glu Gln His
            100                 105                 110
Pro Glu Leu Thr Pro Leu Leu Glu Lys Glu Arg Asp Gly Leu Arg Cys
            115                 120                 125
Arg Gly Asn Arg Ser Pro Val Pro Asp Val Glu Asp Pro Ala Thr Glu
        130                 135                 140
Glu Pro Gly Glu Ser Phe Cys Asx Lys Val Met Arg Trp Phe Gln Ala
145                 150                 155                 160
Met Leu Gln Arg Leu Gln Thr Trp Trp His Gly Val Leu Ala Trp Val
            165                 170                 175
Lys Glu Lys Val Val Ala Leu Val His Ala Val Gln Ala Leu Trp Lys
            180                 185                 190
Gln Phe Gln Ser Phe Cys Cys Ser Leu Ser Glu Leu Phe Met Ser Ser
            195                 200                 205
Phe Gln Ser Tyr Gly Ala Pro Arg Gly Asp Lys Glu Glu Leu Thr Pro
        210                 215                 220
Gln Lys Cys Ser Glu Pro Gln Ser Ser Lys
225                 230
```

<210> 167
<211> 958
<212> DNA
<213> Homo sapiens

<400> 167

```
tctaaagctc agtggagctg ggtcatctca ggccttggct ccttgaactt ttggccgcca 60
tgtgcttccc gaaggtcctc tctgatgaca tgaagaagct gaaggcccga atggtaatgc 120
tcctccctac ttctgctcag gggttggggg cctgggtctc agcgtgtgac actgaggaca 180
ctgtgggaca cctgggaccc tggagggaca aggatccggc cctttggtgc caactctgcc 240
tctcttcaca gcaccaggcc atagaaagat tttatgataa aatgcaaaat gcagaatcag 300
gacgtggaca ggtgatgtcg agcctggcag agctggagga cgacttcaaa gagggctacc 360
tggagacagt ggcggcttat tatgaggagc agcacccaga gctcactcct ctacttgaaa 420
aagaaagaga tggattacgg tgccgaggca acagatcccc tgtcccggat gttgaggatc 480
ccgcaaccga ggagcctggg gagagctttt gtgacaaggt catgagatgg ttccaggcca 540
```

```
tgctgcagcg gctgcagacc tggtggcacg gggttctggc ctgggtgaag gagaaggtgg 600
tggccctggt ccatgcagtg caggccctct ggaaacagtt ccagagtttc tgctgctctc 660
tgtcagagct cttcatgtcc tctttccagt cctacggagc cccacggggg gacaaggagg 720
agctgacacc ccagaagtgc tctgaacccc aatcctcaaa atgaagatac tgacaccacc 780
tttgccctcc ccgtcaccgc gcacccaccc tgacccctcc ctcagctgtc ctgtgccccg 840
ccctctcccg cacactcagt cccctgcct ggcgttcctg ccgcagctct gacctggtgc 900
tgtcgccctg gcatcttaat aaaacctgct tatacttccc tggcagggag ataccatg 958
```

```
<210> 168
<211> 234
<212> PRT
<213> Homo sapiens
```

```
<400> 168
Met Cys Phe Pro Lys Val Leu Ser Asp Asp Met Lys Lys Leu Lys Ala
1               5                   10                  15
Arg Met Val Met Leu Leu Pro Thr Ser Ala Gln Gly Leu Gly Ala Trp
            20                  25                  30
Val Ser Ala Cys Asp Thr Glu Asp Thr Val Gly His Leu Gly Pro Trp
        35                  40                  45
Arg Asp Lys Asp Pro Ala Leu Trp Cys Gln Leu Cys Leu Ser Ser Gln
    50                  55                  60
His Gln Ala Ile Glu Arg Phe Tyr Asp Lys Met Gln Asn Ala Glu Ser
65                  70                  75                  80
Gly Arg Gly Gln Val Met Ser Ser Leu Ala Glu Leu Glu Asp Asp Phe
            85                  90                  95
Lys Glu Gly Tyr Leu Glu Thr Val Ala Ala Tyr Tyr Glu Glu Gln His
            100                 105                 110
Pro Glu Leu Thr Pro Leu Leu Glu Lys Glu Arg Asp Gly Leu Arg Cys
            115                 120                 125
Arg Gly Asn Arg Ser Pro Val Pro Asp Val Glu Asp Pro Ala Thr Glu
            130                 135                 140
Glu Pro Gly Glu Ser Phe Cys Asp Lys Val Met Arg Trp Phe Gln Ala
145                 150                 155                 160
Met Leu Gln Arg Leu Gln Thr Trp Trp His Gly Val Leu Ala Trp Val
                165                 170                 175
Lys Glu Lys Val Val Ala Leu Val His Ala Val Gln Ala Leu Trp Lys
            180                 185                 190
Gln Phe Gln Ser Phe Cys Cys Ser Leu Ser Glu Leu Phe Met Ser Ser
            195                 200                 205
Phe Gln Ser Tyr Gly Ala Pro Arg Gly Asp Lys Glu Glu Leu Thr Pro
    210                 215                 220
Gln Lys Cys Ser Glu Pro Gln Ser Ser Lys
225                 230
```

```
<210> 169
<211> 1005
<212> DNA
<213> Homo sapiens
```

```
<400> 169
tgtgctgcta ttgtgtggat gccgcgcgtg tcttctcttc tttccagaga tggctaacag 60
gggcccgagc tatggcttaa gccgagaggt gcaggagaag atcgagcaga agtatgatgc 120
ggacctggag aacaagctgg tggactggat catcctgcag tgcgccgagg acatagagca 180
cccgcccccc ggcagggccc attttcagaa atggttaatg gacgggacgg tcctgtgcaa 240
gctgataaat agtttatacc caccaggaca agagcccata cccaagatct cagagtcaaa 300
gatggctttt aagcagatgg agcaaatctc ccagttccta aaagctgcgg agacctatgg 360
tgtcagaacc accgacatct ttcagacggt ggatctatgg gaagggaagg acatggcagc 420
```

241

```
tgtgcagagg accctgatgg ctttaggcag cgttgcagtc accaaggatg atggctgcta 480
tcggggagag ccatcctggt ttcacaggaa agcccagcag aatcggagag gctttccga 540
ggagcagctt cgccagggac agaacgtaat aggcctgcag atgggcagca caagggagc 600
ctcccaggcg ggcatgacag ggtacgggat gcccaggcag atcatgttag gacgcggcat 660
cctgcccctg gtagagagga cgaatgttcc acaccatggt ctctacgaaa aagaaatagt 720
tagtcacctt ctgaccttct cctctttctc aaagccttct gtccctggtt tttgcaagtg 780
ctgcatttcc gccgagaatc cgcgttgcct actgctgcca cctcctgttc atttagaact 840
atgcaaagac tccgcttccg ttttcctgag ctcctcgggc cccagagtct ctgtttgatt 900
atttatttat ttatttattt atttgccaaa aattctcctc ttcaacttat agaatgcacc 960
taataaagta attaagtctt gtggaaaaaa aaaaaaaaaa aaaaa          1005
```

```
<210> 170
<211> 282
<212> PRT
<213> Homo sapiens
```

```
<400> 170
Met Ala Asn Arg Gly Pro Ser Tyr Gly Leu Ser Arg Glu Val Gln Glu
  1               5                  10                  15
Lys Ile Glu Gln Lys Tyr Asp Ala Asp Leu Glu Asn Lys Leu Val Asp
             20                  25                  30
Trp Ile Ile Leu Gln Cys Ala Glu Asp Ile Glu His Pro Pro Pro Gly
         35                  40                  45
Arg Ala His Phe Gln Lys Trp Leu Met Asp Gly Thr Val Leu Cys Lys
     50                  55                  60
Leu Ile Asn Ser Leu Tyr Pro Pro Gly Gln Glu Pro Ile Pro Lys Ile
 65                  70                  75                  80
Ser Glu Ser Lys Met Ala Phe Lys Gln Met Glu Gln Ile Ser Gln Phe
                 85                  90                  95
Leu Lys Ala Ala Glu Thr Tyr Gly Val Arg Thr Thr Asp Ile Phe Gln
            100                 105                 110
Thr Val Asp Leu Trp Glu Gly Lys Asp Met Ala Ala Val Gln Arg Thr
            115                 120                 125
Leu Met Ala Leu Gly Ser Val Ala Val Thr Lys Asp Asp Gly Cys Tyr
        130                 135                 140
Arg Gly Glu Pro Ser Trp Phe His Arg Lys Ala Gln Gln Asn Arg Arg
145                 150                 155                 160
Gly Phe Ser Glu Glu Gln Leu Arg Gln Gly Gln Asn Val Ile Gly Leu
                165                 170                 175
Gln Met Gly Ser Asn Lys Gly Ala Ser Gln Ala Gly Met Thr Gly Tyr
            180                 185                 190
Gly Met Pro Arg Gln Ile Met Leu Gly Arg Gly Ile Leu Pro Leu Val
        195                 200                 205
Glu Arg Thr Asn Val Pro His His Gly Leu Tyr Glu Lys Glu Ile Val
    210                 215                 220
Ser His Leu Leu Thr Phe Ser Ser Phe Ser Lys Pro Ser Val Pro Gly
225                 230                 235                 240
Phe Cys Lys Cys Cys Ile Ser Ala Glu Asn Pro Arg Cys Leu Leu Leu
                245                 250                 255
Pro Pro Pro Val His Leu Glu Leu Cys Lys Asp Ser Ala Ser Val Phe
            260                 265                 270
Leu Ser Ser Ser Gly Pro Arg Val Ser Val
        275                 280
```

```
<210> 171
<211> 942
<212> DNA
<213> Homo sapiens
```

<400> 171
```
atgagaattg cagtgatttg cttttgcctc ctaggcatca cctgtgccat accagttaaa 60
caggctgatt ctggaagttc tgaggaaaag cagctttaca acaaataccc agatgctgtg 120
gccacatggc taaaccctga cccatctcag aagcagaatc tcctagcccc acagaatgct 180
gtgtcctctg aagaaaccaa tgactttaaa caagagaccc ttccaagtaa gtccaacgaa 240
agccatgacc acatggatga tatggatgat gaagatgatg atgaccatgt ggacagccag 300
gactccattg actcgaacga ctctgatgat gtagatgaca ctgatgattc tcaccagtct 360
gatgagtctc accattctga tgaatctgat gaactggtca ctgattttcc cacggacctg 420
ccagcaaccg aagttttcac tccagttgtc cccacagtag acacatatga tggccgaggt 480
gatagtgtgg tttatggact gaggtcaaaa tctaagaagt ttcgcagacc tgacatccag 540
taccctgatg ctacagacga gcacatcacc tcacacatgg aaagcgagga gttgaatggt 600
gcatacaagg ccatcccccgt tgcccaggac ctgaacgcgc cttctgattg ggacagccgt 660
gggaaggaca gttatgaaac gagtcagctg gatgaccaga gtgctgaagc ccacagccac 720
aagcagtcca gattatataa gcggaaagct aatgatgaga gcaatgagca ttccgatgtg 780
attgatagtc aggaactttc caaagtcagc cgtgaattcc acagccatga atttcacagc 840
catgaagata tgctggttgt agaccccaaa agtaaggaag aagataaaca cctgaaattt 900
cgtatttctc atgaattaga tagtgcatct tctgaggtca at                        942
```

<210> 172
<211> 314
<212> PRT
<213> Homo sapiens

<400> 172
```
Met Arg Ile Ala Val Ile Cys Phe Cys Leu Leu Gly Ile Thr Cys Ala
1               5                   10                  15
Ile Pro Val Lys Gln Ala Asp Ser Gly Ser Ser Glu Glu Lys Gln Leu
                20                  25                  30
Tyr Asn Lys Tyr Pro Asp Ala Val Ala Thr Trp Leu Asn Pro Asp Pro
            35                  40                  45
Ser Gln Lys Gln Asn Leu Leu Ala Pro Gln Asn Ala Val Ser Ser Glu
        50                  55                  60
Glu Thr Asn Asp Phe Lys Gln Glu Thr Leu Pro Ser Lys Ser Asn Glu
65                  70                  75                  80
Ser His Asp His Met Asp Asp Met Asp Asp Glu Asp Asp Asp Asp His
                85                  90                  95
Val Asp Ser Gln Asp Ser Ile Asp Ser Asn Asp Ser Asp Asp Val Asp
            100                 105                 110
Asp Thr Asp Asp Ser His Gln Ser Asp Glu Ser His His Ser Asp Glu
        115                 120                 125
Ser Asp Glu Leu Val Thr Asp Phe Pro Thr Asp Leu Pro Ala Thr Glu
        130                 135                 140
Val Phe Thr Pro Val Val Pro Thr Val Asp Thr Tyr Asp Gly Arg Gly
145                 150                 155                 160
Asp Ser Val Val Tyr Gly Leu Arg Ser Lys Ser Lys Lys Phe Arg Arg
                165                 170                 175
Pro Asp Ile Gln Tyr Pro Asp Ala Thr Asp Glu His Ile Thr Ser His
            180                 185                 190
Met Glu Ser Glu Glu Leu Asn Gly Ala Tyr Lys Ala Ile Pro Val Ala
        195                 200                 205
Gln Asp Leu Asn Ala Pro Ser Asp Trp Asp Ser Arg Gly Lys Asp Ser
        210                 215                 220
Tyr Glu Thr Ser Gln Leu Asp Asp Gln Ser Ala Glu Ala His Ser His
225                 230                 235                 240
Lys Gln Ser Arg Leu Tyr Lys Arg Lys Ala Asn Asp Glu Ser Asn Glu
                245                 250                 255
His Ser Asp Val Ile Asp Ser Gln Glu Leu Ser Lys Val Ser Arg Glu
            260                 265                 270
```

```
Phe His Ser His Glu Phe His Ser His Glu Asp Met Leu Val Val Asp
        275             280             285
Pro Lys Ser Lys Glu Glu Asp Lys His Leu Lys Phe Arg Ile Ser His
        290             295             300
Glu Leu Asp Ser Ala Ser Ser Glu Val Asn
305             310
```

<210> 173
<211> 1524
<212> DNA
<213> Homo sapiens

<400> 173

```
gcagagcaca gcatcgtcgg gaccagactc gtctcaggcc agttgcagcc ttctcagcca 60
aacgccgacc aaggaaaact cactaccatg agaattgcag tgatttgctt ttgcctccta 120
ggcatcacct gtgccatacc agttaaacag gctgattctg gaagttctga ggaaaagcag 180
ctttacaaca ataccccaga tgctgtggcc acatggctaa accctgaccc atctcagaag 240
cagaatctcc tagccccaca gacccttcca agtaagtcca acgaaagcca tgaccacatg 300
gatgatatgg atgatgaaga tgatgatgac catgtggaca gccaggactc cattgactcg 360
aacgactctg atgatgtaga tgacactgat gattctcacc agtctgatga gtctcaccat 420
tctgatgaat ctgatgaact ggtcactgat tttcccacgg acctgccagc aaccgaagtt 480
ttcactccag ttgtccccac agtagacaca tatgatggcc gaggtgatag tgtggtttat 540
ggactgaggt caaaatctaa gaagtttcgc agacctgaca tccagtaccc tgatgctaca 600
gacgaggaca tcacctcaca catggaaagc gaggagttga atggtgcata caaggccatc 660
cccgttgccc aggacctgaa cgcgccttct gattgggaca ccgtgggaa ggacagttat 720
gaaacgagtc agctggatga ccagagtgct gaaacccaca gccacaagca gtccagatta 780
tataagcgga aagccaatga tgagagcaat gagcattccg atgtgattga tagtcaggaa 840
ctttccaaag tcagccgtga attccacagc catgaatttc acagccatga agatatgctg 900
gttgtagacc ccaaaagtaa ggaagaagat aaacacctga aatttcgtat ttctcatgaa 960
ttagtagtg catcttctga ggtcaattaa aaggagaaaa aatacaattt ctcactttgc 1020
atttagtcaa aagaaaaaat gctttatagc aaaatgaaag agaacatgaa atgcttcttt 1080
ctcagtttat tggttgaatg tgtatctatt tgagtctgga ataactaat gtgtttgata 1140
attagtttag tttgtggctt catggaaact ccctgtaaac taaaagcttc agggttatgt 1200
ctatgttcat tctatagaag aaatgcaaac tatcactgta ttttaatatt tgttattctc 1260
tcatgaatag aaatttatgt agaagcaaac aaaatacttt tacccactta aaaagagaat 1320
ataacatttt atgtcactat aatctttgt tttttaagtt agtgtatatt ttgttgtgat 1380
tatcttttg tggtgtgaat aaatctttta tcttgaatgt aataagaatt tggtggtgtc 1440
aattgcttat ttgtttttccc acggttgtcc agcaattaat aaaacataac ctttttttact 1500
gcctaaaaaa aaaaaaaaaa aaaa                                          1524
```

<210> 174
<211> 300
<212> PRT
<213> Homo sapiens

<400> 174

```
Met Arg Ile Ala Val Ile Cys Phe Cys Leu Leu Gly Ile Thr Cys Ala
1               5               10              15
Ile Pro Val Lys Gln Ala Asp Ser Gly Ser Ser Glu Glu Lys Gln Leu
        20              25              30
Tyr Asn Lys Tyr Pro Asp Ala Val Ala Thr Trp Leu Asn Pro Asp Pro
        35              40              45
Ser Gln Lys Gln Asn Leu Leu Ala Pro Gln Thr Leu Pro Ser Lys Ser
        50              55              60
Asn Glu Ser His Asp His Met Asp Asp Met Asp Asp Glu Asp Asp Asp
65              70              75              80
Asp His Val Asp Ser Gln Asp Ser Ile Asp Ser Asn Asp Ser Asp Asp
                85              90              95
```

```
Val Asp Asp Thr Asp Asp Ser His Gln Ser Asp Glu Ser His His Ser
        100             105             110
Asp Glu Ser Asp Glu Leu Val Thr Asp Phe Pro Thr Asp Leu Pro Ala
        115             120             125
Thr Glu Val Phe Thr Pro Val Val Pro Thr Val Asp Thr Tyr Asp Gly
    130             135             140
Arg Gly Asp Ser Val Val Tyr Gly Leu Arg Ser Lys Ser Lys Lys Phe
145             150             155             160
Arg Arg Pro Asp Ile Gln Tyr Pro Asp Ala Thr Asp Glu Asp Ile Thr
                165             170             175
Ser His Met Glu Ser Glu Glu Leu Asn Gly Ala Tyr Lys Ala Ile Pro
            180             185             190
Val Ala Gln Asp Leu Asn Ala Pro Ser Asp Trp Asp Ser Arg Gly Lys
        195             200             205
Asp Ser Tyr Glu Thr Ser Gln Leu Asp Asp Gln Ser Ala Glu Thr His
    210             215             220
Ser His Lys Gln Ser Arg Leu Tyr Lys Arg Lys Ala Asn Asp Glu Ser
225             230             235             240
Asn Glu His Ser Asp Val Ile Asp Ser Gln Glu Leu Ser Lys Val Ser
                245             250             255
Arg Glu Phe His Ser His Glu Phe His Ser His Glu Asp Met Leu Val
            260             265             270
Val Asp Pro Lys Ser Lys Glu Glu Asp Lys His Leu Lys Phe Arg Ile
        275             280             285
Ser His Glu Leu Asp Ser Ala Ser Ser Glu Val Asn
    290             295             300
```

```
<210> 175
<211> 861
<212> DNA
<213> Homo sapiens

<400> 175
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caggctgatt ctggaagttc tgaggaaaag cagaatgctg tgtcctctga agaaaccaat 120
gactttaaac aagagaccct tccaagtaag tccaacgaaa gccatgacca catggatgat 180
atggatgatg aagatgatga tgaccatgtg gacagccagg actccattga ctcgaacgac 240
tctgatgatg tagatgacac tgatgattct caccagtctg atgagtctca ccattctgat 300
gaatctgatg aactggtcac tgattttccc acggacctgc cagcaaccga gttttcact 360
ccagttgtcc ccacagtaga cacatatgat ggccgaggtg atagtgtggt ttatggactg 420
aggtcaaaat ctaagaagtt tcgcagacct gacatccagt accctgatgc tacagacgag 480
cacatcacct cacacatgga aagcgaggag ttgaatggtg catacaaggc catccccgtt 540
gcccaggacc tgaacgcgcc ttctgattgg gacagccgtg ggaggacag ttatgaaacg 600
agtcagctgg atgaccagag tgctgaagcc cacagccaca gcagtccag attatataag 660
cggaaagcta atgatgagag caatgagcat tccgatgtga ttgatagtca ggaactttcc 720
aaagtcagcc gtgaattcca gccatgaa tttcacagcc atgaagatat gctggttgta 780
gaccccaaaa gtaaggaaga agataaacac ctgaaatttc gtatttctca tgaattagat 840
agtgcatctt ctgaggtcaa t                                          861
```

```
<210> 176
<211> 287
<212> PRT
<213> Homo sapiens

<400> 176
Met Arg Ile Ala Val Ile Cys Phe Cys Leu Leu Gly Ile Thr Cys Ala
1               5               10              15
Ile Pro Val Lys Gln Ala Asp Ser Gly Ser Ser Glu Glu Lys Gln Asn
```

```
              20                  25                  30
Ala Val Ser Ser Glu Glu Thr Asn Asp Phe Lys Gln Glu Thr Leu Pro
          35                  40                  45
Ser Lys Ser Asn Glu Ser His Asp His Met Asp Asp Met Asp Asp Glu
      50                  55                  60
Asp Asp Asp Asp His Val Asp Ser Gln Asp Ser Ile Asp Ser Asn Asp
65                  70                  75                  80
Ser Asp Asp Val Asp Asp Thr Asp Asp Ser His Gln Ser Asp Glu Ser
                  85                  90                  95
His His Ser Asp Glu Ser Asp Glu Leu Val Thr Asp Phe Pro Thr Asp
              100                 105                 110
Leu Pro Ala Thr Glu Val Phe Thr Pro Val Val Pro Thr Val Asp Thr
          115                 120                 125
Tyr Asp Gly Arg Gly Asp Ser Val Val Tyr Gly Leu Arg Ser Lys Ser
      130                 135                 140
Lys Lys Phe Arg Arg Pro Asp Ile Gln Tyr Pro Asp Ala Thr Asp Glu
145                 150                 155                 160
His Ile Thr Ser His Met Glu Ser Glu Glu Leu Asn Gly Ala Tyr Lys
                  165                 170                 175
Ala Ile Pro Val Ala Gln Asp Leu Asn Ala Pro Ser Asp Trp Asp Ser
              180                 185                 190
Arg Gly Lys Asp Ser Tyr Glu Thr Ser Gln Leu Asp Asp Gln Ser Ala
          195                 200                 205
Glu Ala His Ser His Lys Gln Ser Arg Leu Tyr Lys Arg Lys Ala Asn
      210                 215                 220
Asp Glu Ser Asn Glu His Ser Asp Val Ile Asp Ser Gln Glu Leu Ser
225                 230                 235                 240
Lys Val Ser Arg Glu Phe His Ser His Glu Phe His Ser His Glu Asp
                  245                 250                 255
Met Leu Val Val Asp Pro Lys Ser Lys Glu Glu Asp Lys His Leu Lys
              260                 265                 270
Phe Arg Ile Ser His Glu Leu Asp Ser Ala Ser Ser Glu Val Asn
          275                 280                 285
```

<210> 177
<211> 3213
<212> DNA
<213> Homo sapiens

<400> 177

```
agagactcaa gatgattccc tttttaccca tgttttctct actattgctg cttattgtta 60
accctataaa cgccaacaat cattatgaca agatcttggc tcatagtcgt atcaggggtc 120
gggaccaagg cccaaatgtc tgtgcccttc aacagatttt gggcaccaaa aagaaatact 180
tcagcacttg taagaactgg tataaaaagt ccatctgtgg acagaaaacg actgtttat 240
atgaatgttg ccctggttat atgagaatgg aaggaatgaa aggctgccca gcagttttgc 300
ccattgacca tgtttatggc actctgggca tcgtgggagc caccacaacg cagcgctatt 360
ctgacgcctc aaaactgagg gaggagatcg agggaaaggg atccttcact tactttgcac 420
cgagtaatga ggcttgggac aacttggatt ctgatatccg tagaggtttg gagagcaacg 480
tgaatgttga attactgaat gctttacata gtcacatgat taataagaga atgttgacca 540
aggacttaaa aaatggcatg attattcctt caatgtataa caatttgggg cttttcatta 600
accattatcc taatggggtt gtcactgtta attgtgctcg aatcatccat gggaaccaga 660
ttgcaacaaa tggtgttgtc catgtcattg accgtgtgct tacacaaatt ggtacctcaa 720
ttcaagactt cattgaagca gaagatgacc tttcatcttt tagagcagct gccatcacat 780
cggacatatt ggaggccctt ggaagagacg gtcacttcac actctttgct cccaccaatg 840
aggcttttga gaaacttcca cgaggtgtcc tagaaaggtt catgggagac aaagtggctt 900
ccgaagctct tatgaagtac cacatcttaa atactctcca gtgttctgag tctattatgg 960
gaggagcagt ctttgagacg ctggaaggaa atacaattga gataggatgt gacggtgaca 1020
gtataacagt aaatggaatc aaaatggtga acaaaaagga tattgtgaca ataatggtg 1080
```

```
tgatccattt gattgatcag gtcctaattc ctgattctgc caaacaagtt attgagctgg 1140
ctggaaaaca gcaaaccacc ttcacggatc ttgtggccca attaggcttg gcatctgctc 1200
tgaggccaga tggagaatac actttgctgg cacctgtgaa taatgcattt tctgatgata 1260
ctctcagcat ggttcagcgc ctccttaaat taattctgca gaatcacata ttgaaagtaa 1320
aagttggcct taatgagctt tacaacgggc aaatactgga aaccatcgga ggcaaacagc 1380
tcagagtctt cgtatatcgt acagctgtct gcattgaaaa ttcatgcatg gagaaaggga 1440
gtaagcaagg gagaaacggt gcgattcaca tattccgcga gatcatcaag ccagcagaga 1500
aatccctcca tgaaaagtta aaacaagata agcgctttag caccttcctc agcctacttg 1560
aagctgcaga cttgaaagag ctcctgacac aacctggaga ctggacatta tttgtgccaa 1620
ccaatgatgc ttttaaggga atgactagtg aagaaaaga aattctgata cgggacaaaa 1680
atgctcttca aaacatcatt ctttatcacc tgacaccagg agttttcatt ggaaaaggat 1740
ttgaacctgg tgttactaac attttaaaga ccacacaagg aagcaaaatc tttctgaaag 1800
aagtaaatga tacacttctg gtgaatgaat tgaaatcaaa agaatctgac atcatgacaa 1860
caaatggtgt aattcatgtt gtagataaac tcctctatcc agcagacaca cctgttggaa 1920
atgatcaact gctggaaata cttaataaat taatcaaata catccaaatt aagtttgttc 1980
gtggtagcac cttcaaagaa atccccgtga ctgtctatac aactaaaatt ataaccaaag 2040
ttgtggaacc aaaaattaaa gtgattgaag gcagtcttca gcctattatc aaaactgaag 2100
gacccacact aacaaagtc aaaattgaag gtgaacctga attcagactg attaaagaag 2160
gtgaaacaat aactgaagtg atccatggag agccaattat taaaaaatac accaaaatca 2220
ttgatggagt gcctgtggaa ataactgaaa aagagacacg agaagaacga atcattacag 2280
gtcctgaaat aaaatacact aggatttcta ctggaggtgg agaaacagaa gaaactctga 2340
agaaattgtt acaagaagag gtcaccaagg tcaccaaatt cattgaaggt ggtgatggtc 2400
atttatttga agatgaagaa attaaaagac tgcttcaggg agacacaccc gtgaggaagt 2460
tgcaagccaa caaaaaagtt caaggttcta gaagacgatt aagggaaggt cgttctcagt 2520
gaaaatccaa aaaccagaaa aaaatgttta tacaacccta agtcaataac ctgaccttag 2580
aaaattgtga gagccaagtt gacttcagga actgaaacat cagcacaaag aagcaatcat 2640
caaataattc tgaacacaaa tttaatattt ttttttctga atgagaaaca tgagggaaat 2700
tgtggagtta gcctcctgtg gtaaaggaat tgaagaaaat ataacacctt acacccttt 2760
tcatcttgac attaaaagtt ctggctaact ttggaatcca ttagagaaaa atccttgtca 2820
ccagattcat tacaattcaa atcgaagagt tgtgaactgt tatcccattg aaaagaccga 2880
gccttgtatg tatgttatgg atacataaaa tgcacgcaag ccattatctc tccatgggaa 2940
gctaagttat aaaaataggt gcttggtgta caaaacttt tatatcaaaa ggctttgcac 3000
atttctatat gagtgggttt actggtaaat tatgttattt tttacaacta attttgtact 3060
ctcagaatgt ttgtcatatg cttcttgcaa tgcatatttt ttaatctcaa acgtttcaat 3120
aaaaccattt ttcagatata aagagaatta cttcaaattg agtaattcag aaaaactcaa 3180
gatttaagtt aaaaagtggt ttggacttgg gaa                         3213
```

<210> 178
<211> 836
<212> PRT
<213> Homo sapiens

<400> 178

```
Met Ile Pro Phe Leu Pro Met Phe Ser Leu Leu Leu Leu Leu Ile Val
 1               5                  10                  15
Asn Pro Ile Asn Ala Asn Asn His Tyr Asp Lys Ile Leu Ala His Ser
            20                  25                  30
Arg Ile Arg Gly Arg Asp Gln Gly Pro Asn Val Cys Ala Leu Gln Gln
        35                  40                  45
Ile Leu Gly Thr Lys Lys Lys Tyr Phe Ser Thr Cys Lys Asn Trp Tyr
    50                  55                  60
Lys Lys Ser Ile Cys Gly Gln Lys Thr Thr Val Leu Tyr Glu Cys Cys
65                  70                  75                  80
Pro Gly Tyr Met Arg Met Glu Gly Met Lys Gly Cys Pro Ala Val Leu
                85                  90                  95
Pro Ile Asp His Val Tyr Gly Thr Leu Gly Ile Val Gly Ala Thr Thr
            100                 105                 110
Thr Gln Arg Tyr Ser Asp Ala Ser Lys Leu Arg Glu Glu Ile Glu Gly
        115                 120                 125
```

```
Lys Gly Ser Phe Thr Tyr Phe Ala Pro Ser Asn Glu Ala Trp Asp Asn
    130             135             140
Leu Asp Ser Asp Ile Arg Arg Gly Leu Glu Ser Asn Val Asn Val Glu
145             150             155                 160
Leu Leu Asn Ala Leu His Ser His Met Ile Asn Lys Arg Met Leu Thr
                165             170             175
Lys Asp Leu Lys Asn Gly Met Ile Ile Pro Ser Met Tyr Asn Asn Leu
            180             185             190
Gly Leu Phe Ile Asn His Tyr Pro Asn Gly Val Val Thr Val Asn Cys
        195             200             205
Ala Arg Ile Ile His Gly Asn Gln Ile Ala Thr Asn Gly Val Val His
    210             215             220
Val Ile Asp Arg Val Leu Thr Gln Ile Gly Thr Ser Ile Gln Asp Phe
225             230             235                 240
Ile Glu Ala Glu Asp Asp Leu Ser Ser Phe Arg Ala Ala Ala Ile Thr
            245             250             255
Ser Asp Ile Leu Glu Ala Leu Gly Arg Asp Gly His Phe Thr Leu Phe
        260             265             270
Ala Pro Thr Asn Glu Ala Phe Glu Lys Leu Pro Arg Gly Val Leu Glu
        275             280             285
Arg Phe Met Gly Asp Lys Val Ala Ser Glu Ala Leu Met Lys Tyr His
    290             295             300
Ile Leu Asn Thr Leu Gln Cys Ser Glu Ser Ile Met Gly Gly Ala Val
305             310             315                 320
Phe Glu Thr Leu Glu Gly Asn Thr Ile Glu Ile Gly Cys Asp Gly Asp
            325             330             335
Ser Ile Thr Val Asn Gly Ile Lys Met Val Asn Lys Lys Asp Ile Val
        340             345             350
Thr Asn Asn Gly Val Ile His Leu Ile Asp Gln Val Leu Ile Pro Asp
    355             360             365
Ser Ala Lys Gln Val Ile Glu Leu Ala Gly Lys Gln Gln Thr Thr Phe
    370             375             380
Thr Asp Leu Val Ala Gln Leu Gly Leu Ala Ser Ala Leu Arg Pro Asp
385             390             395                 400
Gly Glu Tyr Thr Leu Leu Ala Pro Val Asn Asn Ala Phe Ser Asp Asp
            405             410             415
Thr Leu Ser Met Val Gln Arg Leu Leu Lys Leu Ile Leu Gln Asn His
        420             425             430
Ile Leu Lys Val Lys Val Gly Leu Asn Glu Leu Tyr Asn Gly Gln Ile
        435             440             445
Leu Glu Thr Ile Gly Gly Lys Gln Leu Arg Val Phe Val Tyr Arg Thr
    450             455             460
Ala Val Cys Ile Glu Asn Ser Cys Met Glu Lys Gly Ser Lys Gln Gly
465             470             475                 480
Arg Asn Gly Ala Ile His Ile Phe Arg Glu Ile Ile Lys Pro Ala Glu
            485             490             495
Lys Ser Leu His Glu Lys Leu Lys Gln Asp Lys Arg Phe Ser Thr Phe
        500             505             510
Leu Ser Leu Leu Glu Ala Ala Asp Leu Lys Glu Leu Leu Thr Gln Pro
        515             520             525
Gly Asp Trp Thr Leu Phe Val Pro Thr Asn Asp Ala Phe Lys Gly Met
        530             535             540
Thr Ser Glu Glu Lys Glu Ile Leu Ile Arg Asp Lys Asn Ala Leu Gln
545             550             555                 560
Asn Ile Ile Leu Tyr His Leu Thr Pro Gly Val Phe Ile Gly Lys Gly
                565             570             575
Phe Glu Pro Gly Val Thr Asn Ile Leu Lys Thr Thr Gln Gly Ser Lys
            580             585             590
Ile Phe Leu Lys Glu Val Asn Asp Thr Leu Leu Val Asn Glu Leu Lys
```

```
          595                    600                    605
Ser Lys Glu Ser Asp Ile Met Thr Thr Asn Gly Val Ile His Val Val
    610                 615                    620
Asp Lys Leu Leu Tyr Pro Ala Asp Thr Pro Val Gly Asn Asp Gln Leu
625                     630                    635                 640
Leu Glu Ile Leu Asn Lys Leu Ile Lys Tyr Ile Gln Ile Lys Phe Val
                645                    650                    655
Arg Gly Ser Thr Phe Lys Glu Ile Pro Val Thr Val Tyr Thr Thr Lys
            660                    665                    670
Ile Ile Thr Lys Val Val Glu Pro Lys Ile Lys Val Ile Glu Gly Ser
            675                    680                    685
Leu Gln Pro Ile Ile Lys Thr Glu Gly Pro Thr Leu Thr Lys Val Lys
        690        ·           695                    700
Ile Glu Gly Glu Pro Glu Phe Arg Leu Ile Lys Glu Gly Glu Thr Ile
705                     710                    715                 720
Thr Glu Val Ile His Gly Glu Pro Ile Ile Lys Lys Tyr Thr Lys Ile
                725                    730                    735
Ile Asp Gly Val Pro Val Glu Ile Thr Glu Lys Glu Thr Arg Glu Glu
                740                    745                    750
Arg Ile Ile Thr Gly Pro Glu Ile Lys Tyr Thr Arg Ile Ser Thr Gly
        755                    760                    765
Gly Gly Glu Thr Glu Glu Thr Leu Lys Lys Leu Leu Gln Glu Glu Val
    770                    775                    780
Thr Lys Val Thr Lys Phe Ile Glu Gly Gly Asp Gly His Leu Phe Glu
785                     790                    795                 800
Asp Glu .Glu. Ile Lys Arg Leu Leu Gln Gly Asp Thr Pro Val Arg Lys
            805                    810        .          · 815
Leu Gln ·Ala Asn Lys Lys Val Gln Gly Ser Arg Arg Arg Leu Arg Glu
        820                    825                    830
Gly Arg Ser Gln
        835
```

<210> 179
<211> 3077
<212> DNA
<213> Homo sapiens

<400> 179
```
aacagaactg caacggagag actcaagatg attcccttt  tacccatgtt ttctctacta 60
ttgctgctta ttgttaaccc tataaacgcc aacaatcatt atgacaagat cttggctcat 120
agtcgtatca ggggtcggga ccaaggccca aatgtctgtg cccttcaaca gattttgggc 180
accaaaaaga aatacttcag cacttgtaag aactggtata aaaagtccat ctgtggacag 240
aaaacgactg tttttatatga atgttgccct ggttatatga gaatggaagg aatgaaaggc 300
tgcccagcag ttttgcccat tgaccatgtt tatggcactc tgggcatcgt gggagccacc 360
acaacgcagc gctattctga cgcctcaaaa ctgagggagg agatcgaggg aaagggatcc 420
ttcacttact ttgcaccgag taatgaggct tgggacaact tggattctga tatccgtaga 480
ggtttggaga gcaacgtgaa tgttgaatta ctgaatgctt tacatagtca catgattaat 540
aagagaatgt tgaccaagga cttaaaaat  ggcatgatta ttccttcaat gtataacaat 600
ttggggcttt tcattaacca ttatcctaat ggggttgtca ctgttaattg tgctcgaatc 660
atccatggga accagattgc aacaaatggt gttgtccatg tcattgaccg tgtgcttaca 720
caaattggta cctcaattca agacttcatt gaagcagaag atgacctttc atcttttaga 780
gcagctgcca tcacatcgga catattggag gcccttggaa gagacggtca cttcacactc 840
tttgctccca ccaatgaggc ttttgagaaa cttccacgag gtgtcctaga aaggttcatg 900
ggagacaaag tggcttccga agctcttatg aagtaccaca tcttaaatac tctccagtgt 960
tctgagtcta ttatgggagg agcagtcttt gagacgctgg aaggaaatac aattgagata 1020
ggatgtgacg gtgacagtat aacagtaaat ggaatcaaaa tggtgaacaa aaaggatatt 1080
gtgacaaata tggtgtgat  ccatttgatt gatcaggtcc taattcctga ttctgccaaa 1140
caagttattg agctggctgg aaaacagcaa accaccttca cggatcttgt ggcccaatta 1200
```

```
ggcttggcat ctgctctgag gccagatgga gaatacactt tgctggcacc tgtgaataat 1260
gcattttctg atgatactct cagcatggtt cagcgcctcc ttaaattaat tctgcagaat 1320
cacatattga aagtaaaagt tggccttaat gagctttaca acgggcaaat actggaaacc 1380
atcggaggca aacagctcag agtcttcgta tatcgtacag ctgtctgcat tgaaaattca 1440
tgcatggaga aagggagtaa gcaagggaga aacggtgcga ttcacatatt ccgcgagatc 1500
atcaagccag cagagaaatc cctccatgaa aagttaaaac aagataagcg ctttagcacc 1560
ttcctcagcc tacttgaagc tgcagacttg aaagagctcc tgacacaacc tggagactgg 1620
acattatttg tgccaaccaa tgatgctttt aagggaatga ctagtgaaga aaaagaaatt 1680
ctgatacggg acaaaaatgc tcttcaaaac atcattcttt atcacctgac accaggagtt 1740
ttcattggaa aaggatttga acctggtgtt actaacattt taaagaccac acaaggaagc 1800
aaaatctttc tgaaagaagt aaatgataca cttctggtga atgaattgaa atcaaaagaa 1860
tctgacatca tgacaacaaa tggtgtaatt catgttgtag ataaactcct ctatccagca 1920
gacacacctg ttggaaatga tcaactgctg gaaatactta ataaattaat caaatacatc 1980
caaattaagt ttgttcgtgg tagcacccttc aaagaaatcc ccgtgactgt ctataagcca 2040
attattaaaa aatacaccaa aatcattgat ggagtgcctg tggaaataac tgaaaaagag 2100
acacgagaag aacgaatcat tacaggtcct gaaataaaat acactaggat ttctactgga 2160
ggtggagaaa cagaagaaac tctgaagaaa ttgttacaag aagaggtcac caaggtcacc 2220
aaattcattg aaggtggtga tggtcattta tttgaagatg aagaaattaa aagactgctt 2280
cagggagaca cacccgtgag gaagttgcaa gccaacaaaa agttcaagg ttctagaaga 2340
cgattaaggg aaggtcgttc tcagtgaaaa tccaaaaacc agaaaaaaat gtttatacaa 2400
ccctaagtca ataacctgac cttagaaaat tgtgagagcc aagttgactt caggaactga 2460
aacatcagca caaagaagca atcatcaaat aáttctgaac acaaatttaa tatttttttt 2520
tctgaatgag aaacatgagg gaaattgtgg agttagcctc ctgtggagtt agcctcctgt 2580
ggtaaaggaa ttgaagaaaa tataacacct tacacccttt ttcatcttga cattaaaagt 2640
tctggctaac tttggaatcc attagagaaa aatccttgtc accagattca ttacaattca 2700
aatcgaagag ttgtgaactg ttatcccatt gaaaagaccg agccttgtat gtatgttatg 2760
gatacataaa atgcacgcaa gccattatct ctccatggga agctaagtta taaaaatagg 2820
tgcttggtgt acaaaacttt ttatatcaaa aggctttgca catttctata tgagtgggtt 2880
tactggtaaa ttatgttatt ttttacaact aattttgtac tctcagaatg tttgtcatat 2940
gcttcttgca atgcatattt tttaatctca aacgtttcaa taaaaccatt tttcagatat 3000
aaagagaatt acttcaaatt gagtaattca gaaaaactca agatttaagt taaaaagtgg 3060
tttggacttg ggaacag                                                 3077
```

<210> 180
<211> 779
<212> PRT
<213> Homo sapiens

<400> 180
```
Met Ile Pro Phe Leu Pro Met Phe Ser Leu Leu Leu Leu Leu Ile Val
1               5                   10                  15
Asn Pro Ile Asn Ala Asn Asn His Tyr Asp Lys Ile Leu Ala His Ser
            20                  25                  30
Arg Ile Arg Gly Arg Asp Gln Gly Pro Asn Val Cys Ala Leu Gln Gln
        35                  40                  45
Ile Leu Gly Thr Lys Lys Lys Tyr Phe Ser Thr Cys Lys Asn Trp Tyr
    50                  55                  60
Lys Lys Ser Ile Cys Gly Gln Lys Thr Thr Val Leu Tyr Glu Cys Cys
65                  70                  75                  80
Pro Gly Tyr Met Arg Met Glu Gly Met Lys Gly Cys Pro Ala Val Leu
                85                  90                  95
Pro Ile Asp His Val Tyr Gly Thr Leu Gly Ile Val Gly Ala Thr Thr
            100                 105                 110
Thr Gln Arg Tyr Ser Asp Ala Ser Lys Leu Arg Glu Glu Ile Glu Gly
        115                 120                 125
Lys Gly Ser Phe Thr Tyr Phe Ala Pro Ser Asn Glu Ala Trp Asp Asn
    130                 135                 140
Leu Asp Ser Asp Ile Arg Arg Gly Leu Glu Ser Asn Val Asn Val Glu
145                 150                 155                 160
```

Leu Leu Asn Ala Leu His Ser His Met Ile Asn Lys Arg Met Leu Thr
               165   .   170   175

Lys Asp Leu Lys Asn Gly Met Ile Ile Pro Ser Met Tyr Asn Asn Leu
            180   185   190

Gly Leu Phe Ile Asn His Tyr Pro Asn Gly Val Val Thr Val Asn Cys
        195   200   205

Ala Arg Ile Ile His Gly Asn Gln Ile Ala Thr Asn Gly Val Val His
     210   215   220

Val Ile Asp Arg Val Leu Thr Gln Ile Gly Thr Ser Ile Gln Asp Phe
225   230   235   240

Ile Glu Ala Glu Asp Asp Leu Ser Ser Phe Arg Ala Ala Ala Ile Thr
            245   250   255

Ser Asp Ile Leu Glu Ala Leu Gly Arg Asp Gly His Phe Thr Leu Phe
        260   265   270

Ala Pro Thr Asn Glu Ala Phe Glu Lys Leu Pro Arg Gly Val Leu Glu
        275   280   285

Arg Phe Met Gly Asp Lys Val Ala Ser Glu Ala Leu Met Lys Tyr His
     290   295   300

Ile Leu Asn Thr Leu Gln Cys Ser Glu Ser Ile Met Gly Gly Ala Val
305   310   315   320

Phe Glu Thr Leu Glu Gly Asn Thr Ile Glu Ile Gly Cys Asp Gly Asp
            325   330   335

Ser Ile Thr Val Asn Gly Ile Lys Met Val Asn Lys Lys Asp Ile Val
            340   345   350

Thr Asn Asn Gly Val Ile His Leu Ile Asp Gln Val Leu Ile Pro Asp
        355   360   365

Ser Ala Lys Gln Val Ile Glu Leu Ala Gly Lys Gln Gln Thr Thr Phe
     370   375   380

Thr Asp Leu Val Ala Gln Leu Gly Leu Ala Ser Ala Leu Arg Pro Asp
385   390   395   400

Gly Glu Tyr Thr Leu Leu Ala Pro Val Asn Asn Ala Phe Ser Asp Asp
            405   410   415

Thr Leu Ser Met Val Gln Arg Leu Leu Lys Leu Ile Leu Gln Asn His
            420   425   430

Ile Leu Lys Val Lys Val Gly Leu Asn Glu Leu Tyr Asn Gly Gln Ile
            435   .   440   445

Leu Glu Thr Ile Gly Gly Lys Gln Leu Arg Val Phe Val Tyr Arg Thr
     450   455   460

Ala Val Cys Ile Glu Asn Ser Cys Met Glu Lys Gly Ser Lys Gln Gly
465   470   475   480

Arg Asn Gly Ala Ile His Ile Phe Arg Glu Ile Ile Lys Pro Ala Glu
            485   490   495

Lys Ser Leu His Glu Lys Leu Lys Gln Asp Lys Arg Phe Ser Thr Phe
        500   505   510

Leu Ser Leu Leu Glu Ala Ala Asp Leu Lys Glu Leu Leu Thr Gln Pro
        515   520   525

Gly Asp Trp Thr Leu Phe Val Pro Thr Asn Asp Ala Phe Lys Gly Met
        530   535   540

Thr Ser Glu Glu Lys Glu Ile Leu Ile Arg Asp Lys Asn Ala Leu Gln
545   550   555   560

Asn Ile Ile Leu Tyr His Leu Thr Pro Gly Val Phe Ile Gly Lys Gly
            565   570   575

Phe Glu Pro Gly Val Thr Asn Ile Leu Lys Thr Thr Gln Gly Ser Lys
        580   585   590

Ile Phe Leu Lys Glu Val Asn Asp Thr Leu Leu Val Asn Glu Leu Lys
        595   600   605

Ser Lys Glu Ser Asp Ile Met Thr Thr Asn Gly Val Ile His Val Val
     610   615   620

Asp Lys Leu Leu Tyr Pro Ala Asp Thr Pro Val Gly Asn Asp Gln Leu

```
625                           630                           635                           640
Leu Glu Ile Leu Asn Lys Leu Ile Lys Tyr Ile Gln Ile Lys Phe Val
                    645                           650                           655
Arg Gly Ser Thr Phe Lys Glu Ile Pro Val Thr Val Tyr Lys Pro Ile
                    660                           665                           670
Ile Lys Lys Tyr Thr Lys Ile Ile Asp Gly Val Pro Val Glu Ile Thr
            675                           680                           685
Glu Lys Glu Thr Arg Glu Glu Arg Ile Ile Thr Gly Pro Glu Ile Lys
        690                           695                           700
Tyr Thr Arg Ile Ser Thr Gly Gly Gly Glu Thr Glu Glu Thr Leu Lys
705                           710                           715                           720
Lys Leu Leu Gln Glu Glu Val Thr Lys Val Thr Lys Phe Ile Glu Gly
                725                           730                           735
Gly Asp Gly His Leu Phe Glu Asp Glu Glu Ile Lys Arg Leu Leu Gln
            740                           745                           750
Gly Asp Thr Pro Val Arg Lys Leu Gln Ala Asn Lys Lys Val Gln Gly
            755                           760                           765
Ser Arg Arg Arg Leu Arg Glu Gly Arg Ser Gln
    770                           775
```

```
<210> 181
<211> 2088
<212> DNA
<213> Homo sapiens
```

```
<400> 181
gaattcggca cgagcgcgcg gcgaatctca acgctgcgcc gtctgcgggc gcttccgggc  60
caccagtttc tctgctttcc accctggcgc cccccagccc tggctcccca gctgcgctgc  120
cccgggcgtc cacgccctgc gggcttagcg ggttcagtgg gctcaatctg cgcagcgcca  180
cctccatgtt gaccaagcct ctacaggggc ctcccgcgcc cccgggacc cccacgccgc  240
cgccaggagg caaggatcgg gaagcgttcg aggccgagta tcgactcggc cccctcctgg  300
gtaaggggg ctttggcacc gtcttcgcag acaccgcct cacagatcga ctccaggtgg  360
ccatcaaagt gattccccgg aatcgtgtgc tgggctggtc ccccttgtca gactcagtca  420
catgcccact cgaagtcgca ctgctatgga aagtgggtgc aggtggtggg caccctggcg  480
tgatccgcct gcttgactgg tttgagacac aggaaggctt catgctggtc ctcgagcggc  540
ctttgcccgc ccaggatctc tttgactata tcacagagaa gggcccactg ggtgaaggcc  600
caagccgctg cttctttggc caagtagtgg cagccatcca gcactgccat tcccgtggag  660
ttgtccatcg tgacatcaag gatgagaaca tcctgataga cctacgccgt ggctgtgcca  720
aactcattga ttttggttct ggtgccctgc ttcatgatga accctacact gactttgatg  780
ggacaagggt gtacagcccc ccagagtgga tctctcgaca ccagtaccat gcactccgg  840
ccactgtctg gtcactgggc atcctcctct atgacatggt gtgtgggac attccctttg  900
agagggacca ggagattctg gaagctgagc tccacttccc agcccatgtc tccccagact  960
gctgtgccct aatccgccgg tgcctggccc ccaaaccttc ttcccgaccc tcactggaag 1020
agatcctgct ggacccctgg atgcaaacac cagccgagga tgttaccct caaccccctcc 1080
aaaggaggcc ctgcccttt ggcctggtcc ttgctaccct aagcctggcc tggcctggcc 1140
tggcccccaa tggtcagaag agccatccca tggccatgtc acaggatag atggacattt 1200
gttgacttgg ttttacaggt cattaccagt cattaaagtc cagtattact aaggtaaggg 1260
attgaggatc aggggttaga agacataaac caagtttgcc cagttccctt cccaatccta 1320
caaaggagcc ttcctcccag aacctgtggt ccctgatttt ggaggggaa cttcttgctt 1380
ctcattttgc taaggaagtt tattttggtg aagttgttcc cattttgagc cccgggactc 1440
ttattttgat gatgtgtcac cccacattgg cacctcctac taccaccaca caaacttagt 1500
tcatatgctt ttacttgggc aagggtgctt ccttccaat accccagtag cttttatttt 1560
agtaaaggga cccttcccc tagcctaggg tccatattg ggtcaagctg cttacctgcc 1620
tcagcccagg atttttatt ttggggggagg taatgccctg ttgttacccc aaggcttctt 1680
tttttttttt tttttttttg ggtgagggga ccctactttg ttatcccaag tgctcttatt 1740
ctggtgagaa gaaccttaat tccataattt gggaaggaat ggaagatgga caccaccgga 1800
caccaccaga caataggatg ggatggatgg tttttgggg gatgggctag gggaaataag 1860
gcttgctgtt tgttttcctg gggcgctccc tccaattttg cagattttg caacctcctc 1920
```

```
ctgagccggg attgtccaat tactaaaatg taaataatca cgtattgtgg ggaggggagt 1980
tccaagtgtg ccctcctttt ttttcctgcc tggattattt aaaaagccat gtgtggaaac 2040
ccactattta ataaaagtaa tagaatcaga aaaaaaaaaa aaaaaaa        2088
```

```
<210> 182
<211> 334
<212> PRT
<213> Homo sapiens

<400> 182
Met Leu Thr Lys Pro Leu Gln Gly Pro Pro Ala Pro Pro Gly Thr Pro
1               5                   10                  15
Thr Pro Pro Pro Gly Gly Lys Asp Arg Glu Ala Phe Glu Ala Glu Tyr
            20                  25                  30
Arg Leu Gly Pro Leu Leu Gly Lys Gly Gly Phe Gly Thr Val Phe Ala
        35                  40                  45
Gly His Arg Leu Thr Asp Arg Leu Gln Val Ala Ile Lys Val Ile Pro
    50                  55                  60
Arg Asn Arg Val Leu Gly Trp Ser Pro Leu Ser Asp Ser Val Thr Cys
65                  70                  75                  80
Pro Leu Glu Val Ala Leu Leu Trp Lys Val Gly Ala Gly Gly Gly His
                85                  90                  95
Pro Gly Val Ile Arg Leu Leu Asp Trp Phe Glu Thr Gln Glu Gly Phe
            100                 105                 110
Met Leu Val Leu Glu Arg Pro Leu Pro Ala Gln Asp Leu Phe Asp Tyr
        115                 120                 125
Ile Thr Glu Lys Gly Pro Leu Gly Glu Gly Pro Ser Arg Cys Phe Phe
    130                 135                 140
Gly Gln Val Val Ala Ala Ile Gln His Cys His Ser Arg Gly Val Val
145                 150                 155                 160
His Arg Asp Ile Lys Asp Glu Asn Ile Leu Ile Asp Leu Arg Arg Gly
                165                 170                 175
Cys Ala Lys Leu Ile Asp Phe Gly Ser Gly Ala Leu Leu His Asp Glu
            180                 185                 190
Pro Tyr Thr Asp Phe Asp Gly Thr Arg Val Tyr Ser Pro Pro Glu Trp
        195                 200                 205
Ile Ser Arg His Gln Tyr His Ala Leu Pro Ala Thr Val Trp Ser Leu
    210                 215                 220
Gly Ile Leu Leu Tyr Asp Met Val Cys Gly Asp Ile Pro Phe Glu Arg
225                 230                 235                 240
Asp Gln Glu Ile Leu Glu Ala Glu Leu His Phe Pro Ala His Val Ser
                245                 250                 255
Pro Asp Cys Cys Ala Leu Ile Arg Arg Cys Leu Ala Pro Lys Pro Ser
            260                 265                 270
Ser Arg Pro Ser Leu Glu Glu Ile Leu Leu Asp Pro Trp Met Gln Thr
        275                 280                 285
Pro Ala Glu Asp Val Thr Pro Gln Pro Leu Gln Arg Arg Pro Cys Pro
    290                 295                 300
Phe Gly Leu Val Leu Ala Thr Leu Ser Leu Ala Trp Pro Gly Leu Ala
305                 310                 315                 320
Pro Asn Gly Gln Lys Ser His Pro Met Ala Met Ser Gln Gly
                325                 330


<210> 183
<211> 2304
<212> DNA
<213> Homo sapiens
```

<400> 183
```
gtccccgcag cgccgtcgcg ccctcctgcc gcaggccacc gaggccgccg ccgtctagcg 60
ccccgacctc gccaccatga gagccctgct ggcgcgcctg cttctctgcg tcctggtcgt 120
gagcgactcc aaaggcagca atgaacttca tcaagttcca tcgaactgtg actgtctaaa 180
tggaggaaca tgtgtgtcca acaagtactt ctccaacatt cactggtgca actgcccaaa 240
gaaattcgga gggcagcact gtgaaataga taagtcaaaa acctgctatg aggggaatgg 300
tcactttttac cgaggaaagg ccagcactga caccatgggc cggccctgcc tgccctggaa 360
ctctgccact gtccttcagc aaacgtacca tgcccacaga tctgatgctc ttcagctggg 420
cctgggggaaa cataattact gcaggaaccc agacaaccgg aggcgaccct ggtgctatgt 480
gcaggtgggc ctaaagccgc ttgtccaaga gtgcatggtg catgactgcg cagatggaaa 540
aaagcccctcc tctcctccag aagaattaaa atttcagtgt ggccaaaaga ctctgaggcc 600
ccgctttaag attattgggg gagaattcac caccatcgag aaccagccct ggtttgcggc 660
catctacagg aggcaccggg ggggctctgt cacctacgtg tgtggaggca gcctcatcag 720
cccttgctgg gtgatcagcg ccacacactg cttcattgat tacccaaaga aggaggacta 780
catcgtctac ctgggtcgct caaggcttaa ctccaacacg caaggggaga tgaagtttga 840
ggtggaaaac ctcatcctac acaaggacta cagcgctgac acgcttgctc accacaacga 900
cattgccttg ctgaagatcc gttccaagga gggcaggtgt gcgcagccat cccggactat 960
acagaccatc tgcctgcccc cgatgtataa cgatccccag tttggcacaa gctgtgagat 1020
cactggcttt ggaaaagaga attctaccga ctatctctat ccggagcagc tgaaaatgac 1080
tgttgtgaag ctgatttccc accgggagtg tcagcagccc cactactacg gctctgaagt 1140
caccaccaaa atgctatgtg ctgctgaccc ccaatggaaa acagattcct gccagggaga 1200
ctcaggggga cccctcgtct gttccctcca aggccgcatg actttgactg gaattgtgag 1260
ctggggccgt ggatgtgccc tgaaggacaa gccaggcgtc tacacgagag tctcacactt 1320
cttaccctgg atccgcagtc acaccaagga agagaatggc ctggccctct gagggtcccc 1380
agggaggaaa cgggcaccac ccgctttctt gctggttgtc atttttgcag tagagtcatc 1440
tccatcagct gtaagaagag actgggaaga taggctctgc acagatggat ttgcctgtgg 1500
caccaccagg gtgaacgaca atagctttac cctcacggat aggcctgggt gctggctgcc 1560
cagaccctct ggccaggatg gaggggtggt cctgactcaa catgttactg accagcaact 1620
tgtcttttttc tggactgaag cctgcaggag ttaaaaaggg cagggcatct cctgtgcatg 1680
ggctcgaagg gagagccagc tcccccgacc ggtgggcatt tgtgaggccc atggttgaga 1740
aatgaataat ttcccaatta ggaagtgtaa gcagctgagg tctcttgagg gagcttagcc 1800
aatgtgggag cagcggtttg gggagcagag acactaacga cttcagggca gggctctgat 1860
attccatgaa tgtatcagga aatatatatg tgtgtgtatg tttgcacact tgttgtgtgg 1920
gctgtgagtg taagtgtgag taagagctgg tgtctgattg ttaagtctaa atatttcctt 1980
aaactgtgtg gactgtgatg ccacacagag tggtctttct ggagaggtta taggtcactc 2040
ctggggcctc ttgggtcccc cacgtgacag tgcctgggaa tgtacttatt ctgcagcatg 2100
acctgtgacc agcactgtct cagtttcact ttcatagagt tgtccctttc ttggccagtt 2160
atcccttcct tttagcctag ttcatccaat cctcactggg tggggtgagg accactcctt 2220
acactgaata tttatatttc actatttta tttatatttt tgtaatttta aataaaagtg 2280
atcaataaaa tgtgattttt ctga                                        2304
```

<210> 184
<211> 431
<212> PRT
<213> Homo sapiens

<400> 184
```
Met Arg Ala Leu Leu Ala Arg Leu Leu Leu Cys Val Leu Val Val Ser
1               5                   10                  15
Asp Ser Lys Gly Ser Asn Glu Leu His Gln Val Pro Ser Asn Cys Asp
            20                  25                  30
Cys Leu Asn Gly Gly Thr Cys Val Ser Asn Lys Tyr Phe Ser Asn Ile
        35                  40                  45
His Trp Cys Asn Cys Pro Lys Lys Phe Gly Gly Gln His Cys Glu Ile
    50                  55                  60
Asp Lys Ser Lys Thr Cys Tyr Glu Gly Asn Gly His Phe Tyr Arg Gly
65                  70                  75                  80
Lys Ala Ser Thr Asp Thr Met Gly Arg Pro Cys Leu Pro Trp Asn Ser
                85                  90                  95
```

```
Ala Thr Val Leu Gln Gln Thr Tyr His Ala His Arg Ser Asp Ala Leu
            100             105              110
Gln Leu Gly Leu Gly Lys His Asn Tyr Cys Arg Asn Pro Asp Asn Arg
            115             120             125
Arg Arg Pro Trp Cys Tyr Val Gln Val Gly Leu Lys Pro Leu Val Gln
        130             135             140
Glu Cys Met Val His Asp Cys Ala Asp Gly Lys Lys Pro Ser Ser Pro
145             150             155             160
Pro Glu Glu Leu Lys Phe Gln Cys Gly Gln Lys Thr Leu Arg Pro Arg
                165             170             175
Phe Lys Ile Ile Gly Gly Glu Phe Thr Thr Ile Glu Asn Gln Pro Trp
            180             185             190
Phe Ala Ala Ile Tyr Arg Arg His Arg Gly Gly Ser Val Thr Tyr Val
        195             200             205
Cys Gly Gly Ser Leu Ile Ser Pro Cys Trp Val Ile Ser Ala Thr His
    210             215             220
Cys Phe Ile Asp Tyr Pro Lys Lys Glu Asp Tyr Ile Val Tyr Leu Gly
225             230             235             240
Arg Ser Arg Leu Asn Ser Asn Thr Gln Gly Glu Met Lys Phe Glu Val
            245             250             255
Glu Asn Leu Ile Leu His Lys Asp Tyr Ser Ala Asp Thr Leu Ala His
            260             265             270
His Asn Asp Ile Ala Leu Leu Lys Ile Arg Ser Lys Glu Gly Arg Cys
        275             280             285
Ala Gln Pro Ser Arg Thr Ile Gln Thr Ile Cys Leu Pro Ser Met Tyr
    290             295             300
Asn Asp Pro Gln Phe Gly Thr Ser Cys Glu Ile Thr Gly Phe Gly Lys
305             310             315             320
Glu Asn Ser Thr Asp Tyr Leu Tyr Pro Glu Gln Leu Lys Met Thr Val
            325             330             335
Val Lys Leu Ile Ser His Arg Glu Cys Gln Gln Pro His Tyr Tyr Gly
        340             345             350
Ser Glu Val Thr Thr Lys Met Leu Cys Ala Ala Asp Pro Gln Trp Lys
    355             360             365
Thr Asp Ser Cys Gln Gly Asp Ser Gly Gly Pro Leu Val Cys Ser Leu
370             375             380
Gln Gly Arg Met Thr Leu Thr Gly Ile Val Ser Trp Gly Arg Gly Cys
385             390             395             400
Ala Leu Lys Asp Lys Pro Gly Val Tyr Thr Arg Val Ser His Phe Leu
            405             410             415
Pro Trp Ile Arg Ser His Thr Lys Glu Glu Asn Gly Leu Ala Leu
            420             425             430
```

```
<210> 185
<211> 2123
<212> DNA
<213> Homo sapiens

<400> 185
gggaggagcg gagcggtgcg gaggctctgc tcggatcgag gtctgcagcg cagcttcggg 60
agcatgagtg ctgcagtgac tgcagggaag ctggcacggg caccggccga ccctgggaaa 120
gccggggtcc ccggagttgc agctcccgga gctccggcgg cggctccacc ggcgaaagag 180
atcccggagg tcctagtgga cccacgcagc cggcggcgct atgtgcgggg ccgctttttg 240
ggcaagggcg gctttgccaa gtgcttcgag atctcggacg cggacaccaa ggaggtgttc 300
gcgggcaaga ttgtgcctaa gtctctgctg ctcaagccgc accagaggga agatgtgcc 360
atggaaatat ccattcaccg cagcctcgcc accagcacg tcgtaggatt ccacggcttt 420
ttcgaggaca cgacttcgt gttcgtggtg ttggagctct gccgccggag gtctctcctg 480
gagccgcaca agaggaggaa agccctgact gagcctgagg cccgatacta cctacggcaa 540
```

```
attgtgcttg gctgccagta cctgcaccga aaccgagtta ttcatcgaga cctcaagctg 600
ggcaaccttt tcctgaatga agatctggag gtgaaaatag gggattttgg actggcaacc 660
aaagtcgaat atgacgggga gaggaagaag accctgtgtg ggactcctaa ttacatagct 720
cccgaggtgc tgagcaagaa agagcacagt ttcgaggtgg atgtgtggtc cattgggtgt 780
atcatgtata ccttgttagt gggcaaacca ccttttgaga cttcttgcct aaaagagacc 840
tacctccgga tcaagaagaa tgaatacagt attcccaagc acatcaaccc cgtggccgcc 900
tccctcatcc agaagatgct tcagacagat cccactgccc gcccaaccat taacgagctg 960
cttaatgacg agttctttac ttctggctat atccctgccc gtctccccat cacctgcctg 1020
accattccac caaggttttc gattgctccc agcagcctgg accccagcaa ccggaagccc 1080
ctcacagtcc tcaataaagg cttggagaac cccctgcctg agcgtccccg ggaaaaagaa 1140
gaaccagtgg ttcgagagac aggtgaggtg gtcgactgcc acctcagtga catgctgcag 1200
cagctgcaca gtgtcaatgc ctccaagccc tcggagcgtg ggctggtcag gcaagaggag 1260
gctgaggatc ctgcctgcat ccccatcttc tgggtcagca agtgggtgga ctattcggac 1320
aagtacggcc ttgggtatca gctctgtgat aacagcgtgg gggtgctctt caatgactca 1380
acacgcctca tcctctacaa tgatggtgac agcctgcagt acatagagcg tgacggcact 1440
gagtcctacc tcaccgtgag ttcccatccc aactccttga tgaagaagat caccctcctt 1500
aaatatttcc gcaattacat gagcgagcac ttgctgaagg caggtgccaa catcacgccg 1560
cgcgaaggtg atgagctcgc ccggctgccc tacctacgga cctggttccg cacccgcagc 1620
gccatcatcc tgcacctcag caacggcagc gtgcagatca acttcttcca ggatcacacc 1680
aagctcatct tgtgcccact gatggcagcc gtgacctaca tcgacgagaa gcgggacttc 1740
cgcacatacc gcctgagtct cctggaggag tacggctgct gcaaggagct ggccagccgg 1800
ctccgctacg cccgcactat ggtggacaag ctgctgagct cacgctcggc cagcaaccgt 1860
ctcaaggcct cctaatagct gccctcccct ccggactggt gccctcctca ctcccacctg 1920
catctggggc ccatactggt tggctcccgc ggtgccatgt ctgcagtgtg cccccccagcc 1980
ccggtggctg ggcagagctg catcatcctt gcaggtgggg gttgctgtat aagttatttt 2040
tgtacatgtt cgggtgtggg ttctacagac ttgtccccct ccccctcaac cccaccatat 2100
gaattgtaca gaatatttct att                                          2123
```

```
<210> 186
<211> 603
<212> PRT
<213> Homo sapiens
```

```
<400> 186
Met Ser Ala Ala Val Thr Ala Gly Lys Leu Ala Arg Ala Pro Ala Asp
 1               5                  10                  15
Pro Gly Lys Ala Gly Val Pro Gly Val Ala Ala Pro Gly Ala Pro Ala
            20                  25                  30
Ala Ala Pro Pro Ala Lys Glu Ile Pro Glu Val Leu Val Asp Pro Arg
        35                  40                  45
Ser Arg Arg Arg Tyr Val Arg Gly Arg Phe Leu Gly Lys Gly Gly Phe
    50                  55                  60
Ala Lys Cys Phe Glu Ile Ser Asp Ala Asp Thr Lys Glu Val Phe Ala
65                  70                  75                  80
Gly Lys Ile Val Pro Lys Ser Leu Leu Leu Lys Pro His Gln Arg Glu
                85                  90                  95
Lys Met Ser Met Glu Ile Ser Ile His Arg Ser Leu Ala His Gln His
            100                 105                 110
Val Val Gly Phe His Gly Phe Phe Glu Asp Asn Asp Phe Val Phe Val
        115                 120                 125
Val Leu Glu Leu Cys Arg Arg Arg Ser Leu Leu Glu Pro His Lys Arg
    130                 135                 140
Arg Lys Ala Leu Thr Glu Pro Glu Ala Arg Tyr Tyr Leu Arg Gln Ile
145                 150                 155                 160
Val Leu Gly Cys Gln Tyr Leu His Arg Asn Arg Val Ile His Arg Asp
                165                 170                 175
Leu Lys Leu Gly Asn Leu Phe Leu Asn Glu Asp Leu Glu Val Lys Ile
            180                 185                 190
Gly Asp Phe Gly Leu Ala Thr Lys Val Glu Tyr Asp Gly Glu Arg Lys
```

```
              195                     200                     205
Lys Thr Leu Cys Gly Thr Pro Asn Tyr Ile Ala Pro Glu Val Leu Ser
    210                     215                     220
Lys Lys Glu His Ser Phe Glu Val Asp Val Trp Ser Ile Gly Cys Ile
225                     230                     235                     240
Met Tyr Thr Leu Leu Val Gly Lys Pro Pro Phe Glu Thr Ser Cys Leu
                245                     250                     255
Lys Glu Thr Tyr Leu Arg Ile Lys Lys Asn Glu Tyr Ser Ile Pro Lys
                260                     265                     270
His Ile Asn Pro Val Ala Ala Ser Leu Ile Gln Lys Met Leu Gln Thr
                275                     280                     285
Asp Pro Thr Ala Arg Pro Thr Ile Asn Glu Leu Leu Asn Asp Glu Phe
    290                     295                     300
Phe Thr Ser Gly Tyr Ile Pro Ala Arg Leu Pro Ile Thr Cys Leu Thr
305                     310                     315                     320
Ile Pro Pro Arg Phe Ser Ile Ala Pro Ser Ser Leu Asp Pro Ser Asn
                325                     330                     335
Arg Lys Pro Leu Thr Val Leu Asn Lys Gly Leu Glu Asn Pro Leu Pro
                340                     345                     350
Glu Arg Pro Arg Glu Lys Glu Glu Pro Val Val Arg Glu Thr Gly Glu
                355                     360                     365
Val Val Asp Cys His Leu Ser Asp Met Leu Gln Gln Leu His Ser Val
    370                     375                     380
Asn Ala Ser Lys Pro Ser Glu Arg Gly Leu Val Arg Gln Glu Glu Ala
385                     390                     395                     400
Glu Asp Pro Ala Cys Ile Pro Ile Phe Trp Val Ser Lys Trp Val Asp
                405                     410                     415
Tyr Ser Asp Lys Tyr Gly Leu Gly Tyr Gln Leu Cys Asp Asn Ser Val
    420                     425                     430
Gly Val Leu Phe Asn Asp Ser Thr Arg Leu Ile Leu Tyr Asn Asp Gly
    435                     440                     445
Asp Ser Leu Gln Tyr Ile Glu Arg Asp Gly Thr Glu Ser Tyr Leu Thr
    450                     455                     460
Val Ser Ser His Pro Asn Ser Leu Met Lys Lys Ile Thr Leu Leu Lys
465                     470                     475                     480
Tyr Phe Arg Asn Tyr Met Ser Glu His Leu Leu Lys Ala Gly Ala Asn
                485                     490                     495
Ile Thr Pro Arg Glu Gly Asp Glu Leu Ala Arg Leu Pro Tyr Leu Arg
                500                     505                     510
Thr Trp Phe Arg Thr Arg Ser Ala Ile Ile Leu His Leu Ser Asn Gly
                515                     520                     525
Ser Val Gln Ile Asn Phe Phe Gln Asp His Thr Lys Leu Ile Leu Cys
    530                     535                     540
Pro Leu Met Ala Ala Val Thr Tyr Ile Asp Glu Lys Arg Asp Phe Arg
545                     550                     555                     560
Thr Tyr Arg Leu Ser Leu Leu Glu Glu Tyr Gly Cys Cys Lys Glu Leu
                565                     570                     575
Ala Ser Arg Leu Arg Tyr Ala Arg Thr Met Val Asp Lys Leu Leu Ser
    580                     585                     590
Ser Arg Ser Ala Ser Asn Arg Leu Lys Ala Ser
    595                     600
```

<210> 187
<211> 2617
<212> DNA
<213> Homo sapiens

<400> 187

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aagcagtctc aagcctgccg cagggagaag atggcggtcg ccgtgagaac tttgcaggaa 60
cagctggaaa aggccaaaga gagtcttaag aacgtggatg agaacattcg caagctcacc 120
gggcgggacc cgaatgatgt gaggcccatc caagccagat tgctggccct ttctggtcct 180
ggtggaggta gaggacgtgg tagtttattg ctgaggcgtg gattctcaga tagtggagga 240
cccccagcca aacagagaga ccttgaaggg gcagtcagta ggctgggcgg ggagcgtcgg 300
accagaagag aatcacgcca ggaaagcgac ccggaggatg atgatgttaa aaagccagca 360
ttgcagtctt cagttgtagc tacctccaaa gagcgcacac gtagagacct tatccaggat 420
caaaatatgg atgaaaaggg aaagcaaagg aaccgacgaa tatttggctt attgatgggc 480
actcttcaga aatttaaaca agaatccact gttgctactg aaaggcaaaa caggcgccag 540
gaaattgaac aaaaacttga agtgcaggcg gaagaagaaa gaaagcaggt tgaaaatgaa 600
aggagagaac tgtttgaaga gaggcgtgct aaacagacag aactgcggct tttagaacag 660
aaggttgagc ttgcgcagct gcaagaagaa tggaatgaac ataatgccaa aataattaaa 720
tatataagaa ctaagacaaa gccccatttg ttttatattc ccggaagaat gtgtccagct 780
acccaaaaac taatagaaga gtcacagaga aaaatgaacg ctttatttga tggtagacgc 840
atcgaatttg cagaacaaat aaataaaatg gaggctaggc ctagaagaca atcaatgaag 900
gaaaaagagc atcaggtggt gcgtaatgaa gaacacaagg cggaacaaga agagggtaag 960
gtggctcagc gagaggaaga gttggtggag acaggtaacc agcacaatga tgttgaaata 1020
gaggaagcag gagaggaaga ggaaaaggaa ataggattg ttcatagtga tgcagagaaa 1080
gagcaggagg aggaggaaca aaaacaggaa atggaggtta agatggagga ggaaactgag 1140
gtaagggaaa gtgagaagca gcaggatagt cagcctgaag aagttatgga tgtgctagag 1200
atggttgaga atgtcaaaca tgtaattgct gaccaggagg taatggaaac taatcgagtt 1260
gaaagtgtag aaccttcaga aaatgaagct agcaaagaat tggaaccaga aatggaattt 1320
gaaattgagc cagataaaga atgtaaatcc ctttctcctg ggaaagagaa tgtcagtgct 1380
ttagacatgg aaaaggagtc tgacgaaaaa gaagaaaaag aatctgagcc ccaacctgag 1440
cctgtggctc aacctcaggc tcagtctcag ccccagctcc agcttcaatc ccagtccgag 1500
ccacagcctc agctacaacc tgagcctgct caacctcagc ttcagtctca gccccagctt 1560
cagcttcaat cccagtgcca tgcagtactc cagtcccatc ctccctctca acctgaggat 1620
ttgtcattag ctgtttttaca gccaacaccc caagttactc aggagcatgg gcattttcta 1680
cctgagagga aggattttcc tgtagagtct gtaaaactga ctgaggtacc agtagaccca 1740
gtcttgacag tacatccaga gagcgagagc gaaaccaata ctaggagcag gagtagaggt 1800
cgaactagaa atagaaccac caagagtaga agtcgaagca gtagcagtag cagttctagt 1860
agcagttcaa ccagtagcag cagtggaagt agttccagca gtggaagtag tagcagtcgc 1920
agtagttcca gtagcagctc cagtacaagt ggcagcagca gcagagatag cagcagcagc 1980
actagtagta gtagtgagag tagaagtcgg agtaggggcc ggggacataa tagagataga 2040
aagcacagaa ggagcgtgga tcggaagaga agggatactt caggactaga aagaagtcac 2100
aaatcttcaa aaggtggtag tagtagagat acaaaaggat caaaggataa gaattcccgg 2160
tccgacagaa agaggtctat atcagagagt agtcgatcag gcaaaagatc ttcaagaagt 2220
gaaagagacc gaaaatcaga caggaaagac aaaaggcgtt aatggaagaa gccaggcttt 2280
cttagccatt ctttgcagca gaagatttct tgatgaaaaa ggattacctt ccttgtaaa 2340
gaggatgctg ccttaagaat tgcatgttgt aaaaaatctt tttggaagat acagactgtt 2400
tgtttaccag acattcttgt acttttgca taattttgta agagttattt atcaaaatta 2460
tgtgaggttc caaaatatgt aaaaatgata ataataaaaa aagattaaca tcccttgtca 2520
tcttttttaa atatcctata ctcttcagta agaatctgta tattttaata ggcaaatctt 2580
taagtctgtt cccttcaatt ctgtatcata cattgct                            2617
```

<210> 188
<211> 743
<212> PRT
<213> Homo sapiens

<400> 188
```
Met Ala Val Ala Val Arg Thr Leu Gln Glu Gln Leu Glu Lys Ala Lys
1               5                   10                  15
Glu Ser Leu Lys Asn Val Asp Glu Asn Ile Arg Lys Leu Thr Gly Arg
                20                  25                  30
Asp Pro Asn Asp Val Arg Pro Ile Gln Ala Arg Leu Leu Ala Leu Ser
        35                  40                  45
Gly Pro Gly Gly Gly Arg Gly Arg Gly Ser Leu Leu Leu Arg Arg Gly
    50                  55                  60
```

258

```
Phe Ser Asp Ser Gly Gly Pro Pro Ala Lys Gln Arg Asp Leu Glu Gly
65              70              75              80
Ala Val Ser Arg Leu Gly Gly Glu Arg Arg Thr Arg Arg Glu Ser Arg
            85              90              95
Gln Glu Ser Asp Pro Glu Asp Asp Val Lys Lys Pro Ala Leu Gln
            100             105             110
Ser Ser Val Val Ala Thr Ser Lys Glu Arg Thr Arg Arg Asp Leu Ile
        115             120             125
Gln Asp Gln Asn Met Asp Glu Lys Gly Lys Gln Arg Asn Arg Arg Ile
    130             135             140
Phe Gly Leu Leu Met Gly Thr Leu Gln Lys Phe Lys Gln Glu Ser Thr
145             150             155             160
Val Ala Thr Glu Arg Gln Asn Arg Arg Gln Glu Ile Glu Gln Lys Leu
            165             170             175
Glu Val Gln Ala Glu Glu Glu Arg Lys Gln Val Glu Asn Glu Arg Arg
            180             185             190
Glu Leu Phe Glu Glu Arg Arg Ala Lys Gln Thr Glu Leu Arg Leu Leu
        195             200             205
Glu Gln Lys Val Glu Leu Ala Gln Leu Gln Glu Glu Trp Asn Glu His
    210             215             220
Asn Ala Lys Ile Ile Lys Tyr Ile Arg Thr Lys Thr Lys Pro His Leu
225             230             235             240
Phe Tyr Ile Pro Gly Arg Met Cys Pro Ala Thr Gln Lys Leu Ile Glu
            245             250             255
Glu Ser Gln Arg Lys Met Asn Ala Leu Phe Asp Gly Arg Arg Ile Glu
            260             265             270
Phe Ala Glu Gln Ile Asn Lys Met Glu Ala Arg Pro Arg Arg Gln Ser
    275             280             285
Met Lys Glu Lys Glu His Gln Val Val Arg Asn Glu Glu His Lys Ala
    290             295             300
Glu Gln Glu Glu Gly Lys Val Ala Gln Arg Glu Glu Leu Val Glu
305             310             315             320
Thr Gly Asn Gln His Asn Asp Val Glu Ile Glu Glu Ala Gly Glu Glu
            325             330             335
Glu Glu Lys Glu Ile Gly Ile Val His Ser Asp Ala Glu Lys Glu Gln
        340             345             350
Glu Glu Glu Glu Gln Lys Gln Glu Met Glu Val Lys Met Glu Glu Glu
        355             360             365
Thr Glu Val Arg Glu Ser Glu Lys Gln Gln Asp Ser Gln Pro Glu Glu
    370             375             380
Val Met Asp Val Leu Glu Met Val Glu Asn Val Lys His Val Ile Ala
385             390             395             400
Asp Gln Glu Val Met Glu Thr Asn Arg Val Glu Ser Val Glu Pro Ser
            405             410             415
Glu Asn Glu Ala Ser Lys Glu Leu Glu Pro Glu Met Glu Phe Glu Ile
            420             425             430
Glu Pro Asp Lys Glu Cys Lys Ser Leu Ser Pro Gly Lys Glu Asn Val
        435             440             445
Ser Ala Leu Asp Met Glu Lys Glu Ser Asp Glu Lys Glu Glu Lys Glu
    450             455             460
Ser Glu Pro Gln Pro Glu Pro Val Ala Gln Pro Gln Ala Gln Ser Gln
465             470             475             480
Pro Gln Leu Gln Leu Gln Ser Gln Ser Glu Pro Gln Pro Gln Leu Gln
            485             490             495
Pro Glu Pro Ala Gln Pro Gln Leu Gln Ser Gln Pro Gln Leu Gln Leu
            500             505             510
Gln Ser Gln Cys His Ala Val Leu Gln Ser His Pro Pro Ser Gln Pro
        515             520             525
Glu Asp Leu Ser Leu Ala Val Leu Gln Pro Thr Pro Gln Val Thr Gln
```

259

```
        530                     535                     540
Glu His Gly His Phe Leu Pro Glu Arg Lys Asp Phe Pro Val Glu Ser
545                     550                     555                     560
Val Lys Leu Thr Glu Val Pro Val Asp Pro Val Leu Thr Val His Pro
                        565                     570                     575
Glu Ser Glu Ser Glu Thr Asn Thr Arg Ser Arg Ser Arg Gly Arg Thr
                580                     585                     590
Arg Asn Arg Thr Thr Lys Ser Arg Ser Arg Ser Ser Ser Ser Ser Ser
                595                     600                     605
Ser Ser Ser Ser Ser Thr Ser Ser Ser Ser Gly Ser Ser Ser Ser Ser
        610                     615                     620
Gly Ser Ser Ser Ser Arg Ser Ser Ser Ser Ser Ser Ser Ser Thr Ser
625                     630                     635                     640
Gly Ser Ser Ser Arg Asp Ser Ser Ser Ser Thr Ser Ser Ser Ser Glu
                645                     650                     655
Ser Arg Ser Arg Ser Arg Gly Arg Gly His Asn Arg Asp Arg Lys His
                660                     665                     670
Arg Arg Ser Val Asp Arg Lys Arg Arg Asp Thr Ser Gly Leu Glu Arg
                675                     680                     685
Ser His Lys Ser Ser Lys Gly Gly Ser Ser Arg Asp Thr Lys Gly Ser
        690                     695                     700
Lys Asp Lys Asn Ser Arg Ser Asp Arg Lys Arg Ser Ile Ser Glu Ser
705                     710                     715                     720
Ser Arg Ser Gly Lys Arg Ser Ser Arg Ser Glu Arg Asp Arg Lys Ser
                725                     730                     735
Asp Arg Lys Asp Lys Arg Arg
                740
```

```
<210> 189
<211> 1182
<212> DNA
<213> Homo sapiens

<400> 189
gaattccgct agactaagtt ggtcatgatg cagaagctac tcaaatgcag tcggcttgtc 60
ctggctcttg ccctcatcct ggttctggaa tcctcagttc aaggttatcc tacgcagaga 120
gccaggtacc aatgggtgcg ctgcaatcca gacagtaatt ctgcaaactg ccttgaagaa 180
aaaggaccaa tgttcgaact acttccaggt gaatccaaca agatcccccg tctgaggact 240
gacctttttc caaagacgag aatccaggac ttgaatcgta tcttcccact ttctgaggac 300
tactctggat caggcttcgg ctccggctcc ggctctggat caggatctgg gagtggcttc 360
ctaacggaaa tggaacagga ttaccaacta gtagacgaaa gtgatgcttt ccatgacaac 420
cttaggtctc ttgacaggaa tctgccctca gacagccagg acttgggtca acatggatta 480
gaagaggatt ttatgttata aagaggatt ttcccacctt gacaccaggc aatgtagtta 540
gcatatttta tgtaccatgg ttatatgatt aatcttggga caaagaattt tatagaaatt 600
tttaaacatc tgaaaaagaa gcttaagttt tatcatcctt ttttttctca tgaattctta 660
aaggattatg ctttaatgct gttatctatc ttattgttct tgaaaatacc tgcatttttt 720
ggtatcatgt tcaaccaaca tcattatgaa attaattaga ttcccatggc ataaaatgg 780
ctttaaagaa tatatatata ttttaaagt agcttgagaa gcaaattggc aggtaatatt 840
tcatacctaa attaagactc tgacttggat tgtgaattat aatgatatgc cccttttctt 900
ataaaaacaa aaaaaaaata atgaaacaca gtgaatttgt agagtggggg tatttgacat 960
attttacagg gtggagtgta ctatatacta ttacctttga atgtgtttgc agagctagtg 1020
gatgtgtttg tctacaagta tgattgctgt tacataacac cccaaattaa ctcccaaatt 1080
aaaacacagt tgtgctgtca atacctcata ctgctttacc ttttttttcct ggatatctgt 1140
gtattttcaa atgttactat atattaaagc agaaatataa cc                     1182
```

```
<210> 190
<211> 158
<212> PRT
```

<213> Homo sapiens

<400> 190
```
Met Met Gln Lys Leu Leu Lys Cys Ser Arg Leu Val Leu Ala Leu Ala
1           5               10              15
Leu Ile Leu Val Leu Glu Ser Ser Val Gln Gly Tyr Pro Thr Gln Arg
            20              25              30
Ala Arg Tyr Gln Trp Val Arg Cys Asn Pro Asp Ser Asn Ser Ala Asn
        35              40              45
Cys Leu Glu Glu Lys Gly Pro Met Phe Glu Leu Leu Pro Gly Glu Ser
    50              55              60
Asn Lys Ile Pro Arg Leu Arg Thr Asp Leu Phe Pro Lys Thr Arg Ile
65              70              75              80
Gln Asp Leu Asn Arg Ile Phe Pro Leu Ser Glu Asp Tyr Ser Gly Ser
            85              90              95
Gly Phe Gly Ser Gly Ser Gly Ser Gly Ser Gly Ser Gly Ser Gly Phe
            100             105             110
Leu Thr Glu Met Glu Gln Asp Tyr Gln Leu Val Asp Glu Ser Asp Ala
        115             120             125
Phe His Asp Asn Leu Arg Ser Leu Asp Arg Asn Leu Pro Ser Asp Ser
    130             135             140
Gln Asp Leu Gly Gln His Gly Leu Glu Glu Asp Phe Met Leu
145             150             155
```

<210> 191
<211> 1595
<212> DNA
<213> Homo sapiens

<400> 191
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ccggttcgca aagaagctga cttcagaggg ggaaactttc ttcttttagg aggcggttag 60
ccctgttcca cgaacccagg agaactgctg gccagattaa ttagacattg ctatgggaga 120
cgtgtaaaca cactacttat cattgatgca tatataaaac cattttattt tcgctattat 180
ttcagaggaa gcgcctctga tttgtttctt ttttcccttt ttgctctttc tggctgtgtg 240
gtttggagaa agcacagttg gagtagccgg ttgctaaata agtcccgagc gcgagcggag 300
acgatgcagc ggagactggt tcagcagtgg agcgtcgcgg tgttcctgct gagctacgcg 360
gtgccctcct gcgggcgctc ggtggagggt ctcagccgcc gcctcaaaag agctgtgtct 420
gaacatcagc tcctccatga caaggggaag tccatccaag atttacggcg acgattcttc 480
cttcaccatc tgatcgcaga aatccacaca gctgaaatca gagctacctc ggaggtgtcc 540
cctaactcca agccctctcc caacacaaag aaccaccccg tccgatttgg gtctgatgat 600
gagggcagat acctaactca ggaaactaac aaggtggaga cgtacaaaga gcagccgctc 660
aagacacctg ggaagaaaaa gaaaggcaag cccgggaaac gcaaggagca ggaaaagaaa 720
aaacggcgaa ctcgctctgc ctggttagac tctggagtga ctgggagtgg gctagaaggg 780
gaccacctgt ctgacacctc cacaacgtcg ctggagctcg attcacggta acaggcttct 840
ctggcccgta gcctcagcgg ggtgctctca gctgggtttt ggagcctccc ttctgccttg 900
gcttggacaa acctagaatt ttctcccttt atgtatctct atcgattgtg tagcaattga 960
cagagaataa ctcagaatat tgtctgcctt aaagcagtac ccccctacca cacacacccc 1020
tgtcctccag caccatagag aggcgctaga gcccattcct ctttctccac cgtcacccaa 1080
catcaatcct ttaccactct accaaataat ttcatattca agcttcagaa gctagtgacc 1140
atcttcataa tttgctggag aagtgtattt cttccccttta ctctcacacc tgggcaaact 1200
ttcttcagtg ttttttcattt cttacgttct ttcacttcaa gggagaatat agaagcattt 1260
gatattatct acaaacactg cagaacagca tcatgtcata aacgattctg agccattcac 1320
acttttttatt taattaaatg tatttaatta aatctcaaat ttattttaat gtaaagaact 1380
taaattatgt tttaaacaca tgccttaaat ttgtttaatt aaatttaact ctggtttcta 1440
ccagctcata caaaataaat ggtttctgaa aatgtttaag tattaactta caaggatata 1500
ggttttctc atgtatcttt ttgttcattg gcaagatgaa ataattttc tagggtaatg 1560
ccgtaggaaa aataaaactt cacatttaaa aaaaa                        1595
```

```
<210> 192
<211> 175
<212> PRT
<213> Homo sapiens

<400> 192
Met Gln Arg Arg Leu Val Gln Gln Trp Ser Val Ala Val Phe Leu Leu
1               5                   10                  15
Ser Tyr Ala Val Pro Ser Cys Gly Arg Ser Val Glu Gly Leu Ser Arg
            20                  25                  30·
Arg Leu Lys Arg Ala Val Ser Glu His Gln Leu Leu His Asp Lys Gly
        35                  40                  45
Lys Ser Ile Gln Asp Leu Arg Arg Arg Phe Phe Leu His His Leu Ile
        50                  55                  60
Ala Glu Ile His Thr Ala Glu Ile Arg Ala Thr Ser Glu Val Ser Pro
65                  70                  75                  80
Asn Ser Lys Pro Ser Pro Asn Thr Lys Asn His Pro Val Arg Phe Gly
                85                  90                  95
Ser Asp Asp Glu Gly Arg Tyr Leu Thr Gln Glu Thr Asn Lys Val Glu
            100                 105                 110
Thr Tyr Lys Glu Gln Pro Leu Lys Thr Pro Gly Lys Lys Lys Lys Gly
        115                 120                 125
Lys Pro Gly Lys Arg Lys Glu Gln Glu Lys Lys Lys Arg Arg Thr Arg
    130                 135                 140
Ser Ala Trp Leu Asp Ser Gly Val Thr Gly Ser Gly Leu Glu Gly Asp
145                 150                 155                 160
His Leu Ser Asp Thr Ser Thr Thr Ser Leu Glu Leu Asp Ser Arg
                165                 170                 175


<210> 193
<211> 2657
<212> DNA
<213> Homo sapiens

<220>
<221> misc_feature
<222> 2623, 2624, 2625, 2626, 2627, 2628, 2629
<223> n = A,T,C or G

<400> 193
gaattcggca cgagctgcag ggtcaggagg agaatcgtgg ggccaggagg gcagaggcac 60
actccatctt cgtgctcctc acaggccctg cctccctgcc tgctaaggac acagggaagg 120
gggtccccac ctcagtgcct gcctcccttc cctgtgcctg tgtacctggc agtcacagcc 180
acctggcgtg tcccagaaac caaccggctg acctcatctc ctgcccggcc ccacctccat 240
tggctttggc ttttggcgtt tgtgctgccc gacccctttct cctgtccgga tgcgcagggc 300
agggcctgag ccgtcgagct gcacccacag caggctgcct ttggtgactc accgggtgaa 360
cggggggcatt gcgaggcatc ccctccctgg gtttggctcc tgcccacggg gctgacagta 420
gaaatcacag gctgtgagac agctggagcc cagctctgct tgaacctatt ttaggtctct 480
gatccccgct tcctctttag actcccctag agctcagcca gtgctcaacc tgaggctggg 540
ggtctctgag gaagagtgag ttggagctga ggggtctggg gctgtcccct gagagagggg 600
ccagaggcag tgtcaagagc cgggcagtct gattgtggct caccctccat cactcccagg 660
gcccctggcc cagcagccgc agctcccaac cacaatatcc tttggggtttt ggcctacgga 720
gctggggcgg atgacccca aatagccctg gcagattccc cctagacccg cccgcaccat 780
ggtcaggcat gcccctcctc atcgctggca cagcccagag ggtataaaca gtgctggagg 840
ctggcggggc aggccagctg agtcctgagc agcagcccag cggatcctga gaacttcagg 900
gtgagtttgg ggacccttga ttgttctttc tttttcgcta ttgtaaaatt catgttatat 960
ggagggggca aagtttttcag ggtgttgttt agaatgggaa gatgtcccttt gtatcaccat 1020
ggaccctcat gataatttttg tttctttcac tttctactct gttgacaacc attgtctcct 1080
```

```
cttattttct tttcatttc tgtaacttt tcgttaaact ttagcttgca tttgtaacga 1140
atttttaaat tcacttttgt ttatttgtca gattgtaagt actttctcta atcactttt 1200
tttcaaggca atcagggtat attatattgt acttcagcac agttttagag aacaattgtt 1260
ataattaaat gataaggtag aatatttctg catataaatt ctggctggcg tggaaatatt 1320
cttattggta gaaacaacta catcctggtc atcatcctgc ctttctctt atggttacaa 1380
tgatatacac tgtttgagat gaggataaaa tactctgagt ccaaaccggg cccctctgct 1440
aaccatgttc atgccttctt ctttttccta cagctcctgg gcaacgtgct ggttgttgtg 1500
ctgtctcatc attttggcaa agaattaatt ccaactcaaa aatgcaggct caacagtacc 1560
agcagcagcg tcgaaaattt gcagctgcct tcttggcatt cattttcata ctggcagctg 1620
tggatactgc tgaagcaggg aagaaagaga aaccagaaaa aaagtgaag aagtctgact 1680
gtggagaatg gcagtggagt gtgtgtgtgc ccaccagtgg agactgtggg ctgggcacac 1740
gggagggcac tcggactgga gctgagtgca agcaaaccat gaagacccag agatgtaaga 1800
tcccctgcaa ctggaagaag caatttggcg cggagtgcaa ataccagttc caggcctggg 1860
gagaatgtga cctgaacaca gccctgaaga ccagaactgg aagtctgaag cgagccctgc 1920
acaatgccga atgccagaag actgtcacca tctccaagcc ctgtggcaaa ctgaccaagc 1980
ccaaacctca agcagaatct aagaagaaga aaaggaagg caagaaacag gagaagatgc 2040
tggattaaaa gatgtcacct gtggaacata aaaggacat cagcaaacag gatcaattca 2100
ctcctcaggt gcaggctgcc tatcagaagg tggtggctgg tgtggccaat gccctggctc 2160
acaaatacca ctgagatctt tttccctctg ccaaaaatta tggggacatc atgaagcccc 2220
ttgagcatct gacttctggc taataaagga aatttatttt cattgcaata gtgtgttgga 2280
atttttgtg tctctcactc ggaaggacat atgggaggc aaatcattta aacatcaga 2340
atgagtattt ggtttagagt ttggcaacat atgccatatg ctggctgcca tgaacaaagg 2400
tggctataaa gaggtcatca gtatatgaaa cagccccctg ctgtccattc cttattccat 2460
agaaaagcct tgacttgagg ttagattttt tttatatttt gttttgtgtt attttttct 2520
ttaacatccc taaaatttc cttacatgtt ttactagcca gattttcct cctctcctga 2580
ctactcccag tcatagctgt ccctcttctc ttatgaagat ctnnnnnnnc tcgacctgca 2640
ggcaggcatg caagctt                                             2657
```

<210> 194
<211> 168
<212> PRT
<213> Homo sapiens

<400> 194

```
Met Gln Ala Gln Gln Tyr Gln Gln Gln Arg Arg Lys Phe Ala Ala Ala
 1               5                  10                  15
Phe Leu Ala Phe Ile Phe Ile Leu Ala Ala Val Asp Thr Ala Glu Ala
            20                  25                  30
Gly Lys Lys Glu Lys Pro Glu Lys Lys Val Lys Lys Ser Asp Cys Gly
        35                  40                  45
Glu Trp Gln Trp Ser Val Cys Val Pro Thr Ser Gly Asp Cys Gly Leu
    50                  55                  60
Gly Thr Arg Glu Gly Thr Arg Thr Gly Ala Glu Cys Lys Gln Thr Met
65                  70                  75                  80
Lys Thr Gln Arg Cys Lys Ile Pro Cys Asn Trp Lys Lys Gln Phe Gly
                85                  90                  95
Ala Glu Cys Lys Tyr Gln Phe Gln Ala Trp Gly Glu Cys Asp Leu Asn
            100                 105                 110
Thr Ala Leu Lys Thr Arg Thr Gly Ser Leu Lys Arg Ala Leu His Asn
        115                 120                 125
Ala Glu Cys Gln Lys Thr Val Thr Ile Ser Lys Pro Cys Gly Lys Leu
    130                 135                 140
Thr Lys Pro Lys Pro Gln Ala Glu Ser Lys Lys Lys Lys Lys Glu Gly
145                 150                 155                 160
Lys Lys Gln Glu Lys Met Leu Asp
                165
```

<210> 195

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<211> 2972
<212> DNA
<213> Homo sapiens

<400> 195
tcttcggacc taggctgccc tgccgtcatg tcgcaaggga tcctttctcc gccagcgggc 60
ttgctgtccg atgacgatgt cgtagtttct cccatgtttg agtccacagc tgcagatttg 120
gggtctgtgg tacgcaagaa cctgctatca gactgctctg tcgtctctac ctccctagag 180
gacaagcagc aggttccatc tgaggacagt atggagaagg tgaaagtata cttgagggtt 240
aggcccttgt taccttcaga gttggaacga caggaagatc agggttgtgt ccgtattgag 300
aatgtggaga cccttgttct acaagcaccc aaggactcgt ttgccctgaa gagcaatgaa 360
cggggaattg gccaagccac acacaggttc accttttccc agatctttgg gccagaagtg 420
ggacaggcat ccttcttcaa cctaactgtg aaggagatgg taaaggatgt actcaaaggg 480
cagaactggc tcatctatac atatggagtc actaactcag ggaaaaccca cacgattcaa 540
ggtaccatca aggatggagg gattctcccc cggtccctgg cgctgatctt caatagcctc 600
caaggccaac ttcatccaac acctgatctg aagcccttgc tctccaatga ggtaatctgg 660
ctagacagca agcagatccg acaggaggaa atgaagaagc tgtccctgct aaatggaggc 720
ctccaagagg aggagctgtc cacttccttg aagaggagtg tctacatcga aagtcggata 780
ggtaccagca ccagcttcga cagtggcatt gctgggctct cttctatcag tcagtgtacc 840
agcagtagcc agctggatga aacaagtcat cgatgggcac agccagacac tgccccacta 900
cctgtcccgg caaacattcg cttctccatc tggatctcat tctttgagat ctacaacgaa 960
ctgctttatg acctattaga accgcctagc caacagcgca agaggcagac tttgcggcta 1020
tgcgaggatc aaaatggcaa tccctatgtg aaagatctca actggattca tgtgcaagat 1080
gctgaggagg cctggaagct cctaaaagtg ggtcgtaaga accagagctt tgccagcacc 1140
cacctcaacc agaactccag ccgcagtcac agcatcttct caatcaggat cctacacctt 1200
caggggaag gagatatagt ccccaagatc agcgagctgt cactctgtga tctggctggc 1260
tcagagcgct gcaaagatca gaagagtggt gaacggttga aggaagcagg aaacattaac 1320
acctctctac acaccctggg ccgctgtatt gctgcccttc gtcaaaacca gcagaaccgg 1380
tcaaagcaga acctggttcc cttccgtgac agcaagttga ctcgagtgtt ccaaggtttc 1440
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cctatgcaac tgggattccc atccctgcac tcgttcatca aggaacatag tcttcaggta 1620
tcccccagct tagagaaagg ggctaaggca gacacaggcc ttgatgatga tattgaaaat 1680
gaagctgaca tctccatgta tggcaaagag gagctcctac aagttgtgga agccatgaag 1740
acactgcttt tgaaggaacg acaggaaaag ctacagctgg agatgcatct ccgagatgaa 1800
atttgcaatg agatggtaga acagatgcaa cagcgggaac agtggtgcag tgaacatttg 1860
gacacccaaa aggaactatt ggaggaaatg tatgaagaaa aactaaatat cctcaaggag 1920
tcactgacaa gtttttacca agaagagatt caggagcggg atgaaaagat tgaagagcta 1980
gaagctctct tgcaggaagc cagacaacag tcagtggccc atcagcaatc agggtctgaa 2040
ttggccctac ggcggtcaca aaggttggca gcttctgcct ccacccagca gcttcaggag 2100
gttaaagcta aattacagca gtgcaaagca gagctaaact ctaccactga gagttgcat 2160
aagtatcaga aaatgttaga accaccaccc tcagccaagc ccttcaccat tgatgtggac 2220
aagaagttag aagagggcca gaagaatata aggctgttgc ggacagagct tcagaaactt 2280
ggtgagtctc tccaatcagc agagagagct tgttgccaca gcactggggc aggaaaactt 2340
cgtcaagcct tgaccacttg tgatgacatc ttaatcaaac aggaccagac tctggctgaa 2400
ctgcagaaca acatggtgct agtgaaactg gaccttcgga agaaggcagc atgtattgct 2460
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accaaccagg aaaatcagca accaaaccaa caaccaccag gaagaaacc attccttcga 2580
aatttacttc cccgaacacc aacctgccaa agctcaacag actgcagccc ttatgcccgg 2640
atcctacgct cacggcgttc cccctttactc aaatctgggc cttttggcaa aaagtactaa 2700
ggctgtgggg aaagagaaga gcagtcatgg ccctgaggtg ggtcagctac tctcctgaag 2760
aaataggtct ctttttatgct ttaccatata tcaggaatta tatccaggat gcaatactca 2820
gacactagct tttttctcac ttttgtatta taaccaccta tgtaatctca tgttgttgtt 2880
tttttttatt tacttatatg atttctatgc acacaaaaac agttatatta aagatattat 2940
tgttcacatt ttttattgaa aaaaaaaaa aa                               2972

<210> 196
<211> 890
<212> PRT
```

<213> Homo sapiens

<400> 196
Met Ser Gln Gly Ile Leu Ser Pro Pro Ala Gly Leu Leu Ser Asp Asp
1   5   10   15
Asp Val Val Val Ser Pro Met Phe Glu Ser Thr Ala Ala Asp Leu Gly
            20   25   30
Ser Val Val Arg Lys Asn Leu Leu Ser Asp Cys Ser Val Val Ser Thr
        35   40   45
Ser Leu Glu Asp Lys Gln Gln Val Pro Ser Glu Asp Ser Met Glu Lys
    50   55   60
Val Lys Val Tyr Leu Arg Val Arg Pro Leu Leu Pro Ser Glu Leu Glu
65   70   75   80
Arg Gln Glu Asp Gln Gly Cys Val Arg Ile Glu Asn Val Glu Thr Leu
            85   90   95
Val Leu Gln Ala Pro Lys Asp Ser Phe Ala Leu Lys Ser Asn Glu Arg
            100   105   110
Gly Ile Gly Gln Ala Thr His Arg Phe Thr Phe Ser Gln Ile Phe Gly
            115   120   125
Pro Glu Val Gly Gln Ala Ser Phe Phe Asn Leu Thr Val Lys Glu Met
    130   135   140
Val Lys Asp Val Leu Lys Gly Gln Asn Trp Leu Ile Tyr Thr Tyr Gly
145   150   155   160
Val Thr Asn Ser Gly Lys Thr His Thr Ile Gln Gly Thr Ile Lys Asp
            165   170   175
Gly Gly Ile Leu Pro Arg Ser Leu Ala Leu Ile Phe Asn Ser Leu Gln
            180   185   190
Gly Gln Leu His Pro Thr Pro Asp Leu Lys Pro Leu Leu Ser Asn Glu
            195   200   205
Val Ile Trp Leu Asp Ser Lys Gln Ile Arg Gln Glu Glu Met Lys Lys
    210   215   220
Leu Ser Leu Leu Asn Gly Gly Leu Gln Glu Glu Glu Leu Ser Thr Ser
225   230   235   240
Leu Lys Arg Ser Val Tyr Ile Glu Ser Arg Ile Gly Thr Ser Thr Ser
            245   250   255
Phe Asp Ser Gly Ile Ala Gly Leu Ser Ser Ile Ser Gln Cys Thr Ser
            260   265   270
Ser Ser Gln Leu Asp Glu Thr Ser His Arg Trp Ala Gln Pro Asp Thr
    275   280   285
Ala Pro Leu Pro Val Pro Ala Asn Ile Arg Phe Ser Ile Trp Ile Ser
    290   295   300
Phe Phe Glu Ile Tyr Asn Glu Leu Leu Tyr Asp Leu Leu Glu Pro Pro
305   310   315   320
Ser Gln Gln Arg Lys Arg Gln Thr Leu Arg Leu Cys Glu Asp Gln Asn
            325   330   335
Gly Asn Pro Tyr Val Lys Asp Leu Asn Trp Ile His Val Gln Asp Ala
            340   345   350
Glu Glu Ala Trp Lys Leu Leu Lys Val Gly Arg Lys Asn Gln Ser Phe
            355   360   365
Ala Ser Thr His Leu Asn Gln Asn Ser Ser Arg Ser His Ser Ile Phe
    370   375   380
Ser Ile Arg Ile Leu His Leu Gln Gly Glu Gly Asp Ile Val Pro Lys
385   390   395   400
Ile Ser Glu Leu Ser Leu Cys Asp Leu Ala Gly Ser Glu Arg Cys Lys
            405   410   415
Asp Gln Lys Ser Gly Glu Arg Leu Lys Glu Ala Gly Asn Ile Asn Thr
            420   425   430
Ser Leu His Thr Leu Gly Arg Cys Ile Ala Ala Leu Arg Gln Asn Gln
            435   440   445

```
Gln Asn Arg Ser Lys Gln Asn Leu Val Pro Phe Arg Asp Ser Lys Leu
    450                 455             460
Thr Arg Val Phe Gln Gly Phe Phe Thr Gly Arg Gly Arg Ser Cys Met
465                 470             475             480
Ile Val Asn Val Asn Pro Cys Ala Ser Thr Tyr Asp Glu Thr Leu His
                485             490             495
Val Ala Lys Phe Ser Ala Ile Ala Ser Gln Leu Val His Ala Pro Pro
            500             505             510
Met Gln Leu Gly Phe Pro Ser Leu His Ser Phe Ile Lys Glu His Ser
        515             520             525
Leu Gln Val Ser Pro Ser Leu Glu Lys Gly Ala Lys Ala Asp Thr Gly
    530             535             540
Leu Asp Asp Asp Ile Glu Asn Glu Ala Asp Ile Ser Met Tyr Gly Lys
545             550             555             560
Glu Glu Leu Leu Gln Val Val Glu Ala Met Lys Thr Leu Leu Leu Lys
            565             570             575
Glu Arg Gln Glu Lys Leu Gln Leu Glu Met His Leu Arg Asp Glu Ile
        580             585             590
Cys Asn Glu Met Val Glu Gln Met Gln Gln Arg Glu Gln Trp Cys Ser
    595             600             605
Glu His Leu Asp Thr Gln Lys Glu Leu Leu Glu Glu Met Tyr Glu Glu
    610             615             620
Lys Leu Asn Ile Leu Lys Glu Ser Leu Thr Ser Phe Tyr Gln Glu Glu
625             630             635             640
Ile Gln Glu Arg Asp Glu Lys Ile Glu Glu Leu Glu Ala Leu Leu Gln
            645             650             655
Glu Ala Arg Gln Gln Ser Val Ala His Gln Gln Ser Gly Ser Glu Leu
        660             665             670
Ala Leu Arg Arg Ser Gln Arg Leu Ala Ala Ser Ala Ser Thr Gln Gln
    675             680             685
Leu Gln Glu Val Lys Ala Lys Leu Gln Gln Cys Lys Ala Glu Leu Asn
    690             695             700
Ser Thr Thr Glu Glu Leu His Lys Tyr Gln Lys Met Leu Glu Pro Pro
705             710             715             720
Pro Ser Ala Lys Pro Phe Thr Ile Asp Val Asp Lys Lys Leu Glu Glu
            725             730             735
Gly Gln Lys Asn Ile Arg Leu Leu Arg Thr Glu Leu Gln Lys Leu Gly
            740             745             750
Glu Ser Leu Gln Ser Ala Glu Arg Ala Cys Cys His Ser Thr Gly Ala
        755             760             765
Gly Lys Leu Arg Gln Ala Leu Thr Thr Cys Asp Asp Ile Leu Ile Lys
    770             775             780
Gln Asp Gln Thr Leu Ala Glu Leu Gln Asn Asn Met Val Leu Val Lys
785             790             795             800
Leu Asp Leu Arg Lys Lys Ala Ala Cys Ile Ala Glu Gln Tyr His Thr
            805             810             815
Val Leu Lys Leu Gln Gly Gln Val Ser Ala Lys Lys Arg Leu Gly Thr
        820             825             830
Asn Gln Glu Asn Gln Gln Pro Asn Gln Gln Pro Pro Gly Lys Lys Pro
    835             840             845
Phe Leu Arg Asn Leu Leu Pro Arg Thr Pro Thr Cys Gln Ser Ser Thr
    850             855             860
Asp Cys Ser Pro Tyr Ala Arg Ile Leu Arg Ser Arg Arg Ser Pro Leu
865             870             875             880
Leu Lys Ser Gly Pro Phe Gly Lys Lys Tyr
            885             890
```

<210> 197

```
<211> 768
<212> DNA
<213> Homo sapiens

<400> 197
ccttcagcat aaaagctgat ccacaaacaa gaggagcacc agacctcctc ttggcttcga 60
gatggcttcg ccacaccaag agcccaaacc tggagacctg attgagattt tccgccttgg 120
ctatgagcac tgggccctgt atataggaga tggctacgtg atccatctgg ctcctccaag 180
tgagtacccc ggggctggct cctccagtgt cttctcagtc ctgagcaaca gtgcagaggt 240
gaaacggggg cgcctggaag atgtggtggg aggctgttgc tatcgggtca acaacagctt 300
ggaccatgag taccaaccac ggcccgtgga ggtgatcatc agttctgcga aggagatggt 360
tggtcagaag atgaagtaca gtattgtgag caggaactgt gagcactttg tcgcccagct 420
gagatatggc aagtcccgct gtaaacaggt ggaaaaggcc aaggttgaag tcggtgtggc 480
cacggcgctt ggaatcctgg ttgttgctgg atgctctttt gcgattagga gataccaaaa 540
aaaagcaaca gcctgaagca gccacaaaat cctgtgttag aagcagctgt gggggtccca 600
gtggagatga gcctccccca tgcctccagc agcctgaccc tcgtgccctg tctcaggcgt 660
tctctagatc ctttcctctg tttccctctc tcgctggcaa aagtatgatc taattgaaac 720
aagactgaag gatcaataaa cagccatctg ccccttcaaa aaaaaaaa              768


<210> 198
<211> 164
<212> PRT
<213> Homo sapiens

<400> 198
Met Ala Ser Pro His Gln Glu Pro Lys Pro Gly Asp Leu Ile Glu Ile
1               5                   10                  15
Phe Arg Leu Gly Tyr Glu His Trp Ala Leu Tyr Ile Gly Asp Gly Tyr
                20                  25                  30
Val Ile His Leu Ala Pro Pro Ser Glu Tyr Pro Gly Ala Gly Ser Ser
            35                  40                  45
Ser Val Phe Ser Val Leu Ser Asn Ser Ala Glu Val Lys Arg Gly Arg
    50                  55                  60
Leu Glu Asp Val Val Gly Gly Cys Cys Tyr Arg Val Asn Asn Ser Leu
65                  70                  75                  80
Asp His Glu Tyr Gln Pro Arg Pro Val Glu Val Ile Ile Ser Ser Ala
                85                  90                  95
Lys Glu Met Val Gly Gln Lys Met Lys Tyr Ser Ile Val Ser Arg Asn
            100                 105                 110
Cys Glu His Phe Val Ala Gln Leu Arg Tyr Gly Lys Ser Arg Cys Lys
        115                 120                 125
Gln Val Glu Lys Ala Lys Val Glu Val Gly Val Ala Thr Ala Leu Gly
    130                 135                 140
Ile Leu Val Val Ala Gly Cys Ser Phe Ala Ile Arg Arg Tyr Gln Lys
145                 150                 155                 160
Lys Ala Thr Ala


<210> 199
<211> 720
<212> DNA
<213> Homo sapiens

<400> 199
ggggggggggc ggagggcgct catttccggg ccgcccacca cccgcgtagc accggcagcc 60
gctgtcccgg cagtctccag ccgtcccgcc cgcttgtggc caaactggct ccagtcactc 120
ccgaaatgcc agtcgacttc actgggtact ggaagatgtt ggtcaacgag aatttcgagg 180
agtacctgcg cgccctcgac gtcaatgtgg ccttgcgcaa aatcgccaac ttgctgaagc 240
```

```
cagacaaaga gatcgtgcag gacggtgacc atatgatcat ccgcacgctg agcactttta 300
ggaactacat catggacttc caagttggga aggagtttga ggaggatctg acaggcatag 360
atgaccgcaa gtgcatgaca acagtgagct gggacggaga caagctccag tgtgtgcaga 420
agggtgagaa ggaggggcgt ggctggaccc agtggatcga gggtgatgag ctgcacctag 480
agatgagagt ggaaggtgtg gtctgcaagc aagtattcaa gaaggtgcag tgaggcccaa 540
gcagacaacc ttgtcccaac caatcagcag gatgtgtgag ccaggatccc tctttgcaca 600
gcatgaggca aaaatgtcca gccacccta ggcatctgtt agcagagtct gtctcttggc 660
tttgtcactt ttcctttct taaaacaaag ccatgccaat aaagtgacct gtgttcaaaa 720
```

<210> 200
<211> 135
<212> PRT
<213> Homo sapiens

<400> 200

```
Met Pro Val Asp Phe Thr Gly Tyr Trp Lys Met Leu Val Asn Glu Asn
 1               5                  10                  15
Phe Glu Glu Tyr Leu Arg Ala Leu Asp Val Asn Val Ala Leu Arg Lys
                20                  25                  30
Ile Ala Asn Leu Leu Lys Pro Asp Lys Glu Ile Val Gln Asp Gly Asp
            35                  40                  45
His Met Ile Ile Arg Thr Leu Ser Thr Phe Arg Asn Tyr Ile Met Asp
     50                  55                  60
Phe Gln Val Gly Lys Glu Phe Glu Glu Asp Leu Thr Gly Ile Asp Asp
65                  70                  75                  80
Arg Lys Cys Met Thr Thr Val Ser Trp Asp Gly Asp Lys Leu Gln Cys
                85                  90                  95
Val Gln Lys Gly Glu Lys Glu Gly Arg Gly Trp Thr Gln Trp Ile Glu
            100                 105                 110
Gly Asp Glu Leu His Leu Glu Met Arg Val Glu Gly Val Val Cys Lys
        115                 120                 125
Gln Val Phe Lys Lys Val Gln
        130                 135
```

<210> 201
<211> 2383
<212> DNA
<213> Homo sapiens

<400> 201

```
ggggctaccg cgcctttgct tcctggcgca cgcggagcct cctggagcct gccaccatcc 60
tgcctactac gtgctgccct gcgccgcag ccatgtgccg caccctggcc gccttcccca 120
ccacctgcct ggagagagcc aaagagttca agacacgtct ggggatcttt cttcacaaat 180
cagagctggg ctgcgatact gggagtactg gcaagtccga gtggggcagt aaacacagca 240
aagagaatag aaacttctca gaagatgtgc tggggtggag agagtcgttc gacctgctgc 300
tgagcagtaa aaatggagtg gctgccttcc acgctttcct gaagacagag ttcagtgagg 360
agaacctgga gttctggctg gcctgtgagg agttcaagaa gatccgatca gctaccaagc 420
tggcctccag ggcacaccag atctttgagg agttcatttg cagtgaggcc cctaaagagg 480
tcaacattga ccatgagacc cgcgagctga cgaggatgaa cctgcagact gccacagcca 540
catgctttga tgcggctcag gggaagacac gtaccctgat ggagaaggac tcctacccac 600
gcttcctgaa gtcgcctgct taccgggacc tggctgccca agcctcagcc gcctctgcca 660
ctctgtccag ctgcagcctg gacgagccct cacacacctg agtctccacg gcagtgagga 720
agccagccgg gaagagaggt tgagtcaccc atccccgagg tggctgcccc tgtgtgggag 780
gcaggttctg caaagcaagt gcaagaggac aaaaaaaaaa aaaaaaaaa aaaaatgcg 840
ctccagcagc ctgtttggga agcagcagtc tctccttcag atactgtggg actcatgctg 900
gagaggagcc gcccacttcc aggacctgtg aataagggct aatgatgagg gttggtgggg 960
ctctctgtgg ggcaaaaagg tggtatgggg gttagcactg gctctcgttc tcaccggaga 1020
```

```
aggaagtgtt ctagtgtggt ttaggaaaca tgtggataaa gggaaccatg aaaatgagag 1080
gaggaaagac atccagatca gctgtttttgc ctgttgctca gttgactctg attgcatcct 1140
gttttcctaa ttcccagact gttctgggca cggaagggac cctggatgtg gagtcttccc 1200
ctttggcccт cctcactggc ctctgggcta gcccagagtc ccttagcttg tacctcgtaa 1260
cactcctgtg tgtctgtcca gccttgcagt catgtcaagg ccagcaagct gatgtgactc 1320
ttgccccatg cgagatattt atacctcaaa cactggcctg tgagcccttt ccaagtcagt 1380
ggagagccct gaaaggaggc tcacttgaat ccagctcagt gctctgggtg ccccctgca 1440
ggtggcccct gacctgcgt tgcagcaggg tccacctgtg agcaggcccg ccctggggcc 1500
tcttcctgga tgtgccctct ctgagttctg tgctgtctct tggaggcagg gcccaggaga 1560
acaaagtgtg gaggcctcgg ggagtggctt ttccagctct catgccccgc agtgtggaac 1620
aaggcagaaa aggatcctag gaaataagtc tcttggcggt ccctgagagt cctgctgaaa 1680
tccagccagt gttttttgtg gtatgagaac aggcaaaaag agatgccccg agatagaagg 1740
ggagccttgt gtttctttcc tgcagacgtg agatgaacac tggagtgggc agaggtggcm 1800
caggaccatg gcaccttag agtgcagaag ctgggggagg aggctgcttc gaagggcagg 1860
actggggata cctgcctgtc acctcagggc atcactgaac aaacatttcc tgatggsaac 1920
tcctgcggca gagcccaggc tggggaagtg aactacccag ggcagcccct ttgtggccca 1980
ggataatcaa cactgttctc tctgtaccat gagctcctcc aggagattat ttaagtgtat 2040
tgtatcattg gttttctgtg attgtcataa cattgttttt gttattgttg gtgctgttgt 2100
tatttattat tgtaatttca gtttgcctct actggagaat ctcagcaggg gtttcagcct 2160
gactgtctcc ctttctctac cagactctac ctctgaatgt gctgggaacc tcttggagcc 2220
tgtcaggaac tcctcactgt ttaaatattt atttattgtg acaaatggag ctggtttcct 2280
agatatgaat gatgtttgca atccccattt tcctgtttca gcatgttata ttcttataaa 2340
ataaaagcaa aagtcaaata tgaaaaaaaa aaaaaaaaa aaa 2383
```

<210> 202
<211> 202
<212> PRT
<213> Homo sapiens

<400> 202

```
Met Cys Arg Thr Leu Ala Ala Phe Pro Thr Thr Cys Leu Glu Arg Ala
1               5                   10                  15
Lys Glu Phe Lys Thr Arg Leu Gly Ile Phe Leu His Lys Ser Glu Leu
            20                  25                  30
Gly Cys Asp Thr Gly Ser Thr Gly Lys Ser Glu Trp Gly Ser Lys His
            35                  40                  45
Ser Lys Glu Asn Arg Asn Phe Ser Glu Asp Val Leu Gly Trp Arg Glu
        50                  55                  60
Ser Phe Asp Leu Leu Leu Ser Ser Lys Asn Gly Val Ala Ala Phe His
65                  70                  75                  80
Ala Phe Leu Lys Thr Glu Phe Ser Glu Glu Asn Leu Glu Phe Trp Leu
                85                  90                  95
Ala Cys Glu Glu Phe Lys Lys Ile Arg Ser Ala Thr Lys Leu Ala Ser
            100                 105                 110
Arg Ala His Gln Ile Phe Glu Glu Phe Ile Cys Ser Glu Ala Pro Lys
        115                 120                 125
Glu Val Asn Ile Asp His Glu Thr Arg Glu Leu Thr Arg Met Asn Leu
    130                 135                 140
Gln Thr Ala Thr Ala Thr Cys Phe Asp Ala Ala Gln Gly Lys Thr Arg
145                 150                 155                 160
Thr Leu Met Glu Lys Asp Ser Tyr Pro Arg Phe Leu Lys Ser Pro Ala
                165                 170                 175
Tyr Arg Asp Leu Ala Ala Gln Ala Ser Ala Ala Ser Ala Thr Leu Ser
            180                 185                 190
Ser Cys Ser Leu Asp Glu Pro Ser His Thr
            195                 200
```

<210> 203

```
<211> 616
<212> DNA
<213> Homo sapiens

<400> 203
ctcccctggg agcctggctg ccttgctctc cttcctgggt ctgtctctgc cacctggtct 60
gccacagatc catgatgtgc agttctctgg agcaggcgct ggctgtgctg gtcactacct 120
tccacaagta ctcctgccaa gagggcgaca agttcaagct gagtaagggg gaaatgaagg 180
aacttctgca caaggagctg cccagctttg tggggcattc cagagaacca tgtgctgtga 240
gggccttccg agtccatctg tttaatcctg tcattggaga cttgagaaac cagagcccag 300
aagggaaaag tgattgtccc aagatcacac agcactggag aaagtggatg aggaggggct 360
gaagaagctg atgggcagcc tggatgagaa cagtgaccag caggtggact ccaggagta 420
tgctgttttc ctggcactca tcactgtcat gtgcaatgac ttcttccagg gctgcccaga 480
ccgaccctga agcagaactc ttgacttcct gccatggatc tcttgggccc aggactgttg 540
atgcctttga gttttgtatt caataaactt tttttgtctg ttgaaaaaaa aaaaaaaaaa 600
aaaaaaaaaa aaaaaa                                                 616


<210> 204
<211> 96
<212> PRT
<213> Homo sapiens

<400> 204
Met Met Cys Ser Ser Leu Glu Gln Ala Leu Ala Val Leu Val Thr Thr
1               5                   10                  15
Phe His Lys Tyr Ser Cys Gln Glu Gly Asp Lys Phe Lys Leu Ser Lys
                20                  25                  30
Gly Glu Met Lys Glu Leu Leu His Lys Glu Leu Pro Ser Phe Val Gly
            35                  40                  45
His Ser Arg Glu Pro Cys Ala Val Arg Ala Phe Arg Val His Leu Phe
        50                  55                  60
Asn Pro Val Ile Gly Asp Leu Arg Asn Gln Ser Pro Glu Gly Lys Ser
65                  70                  75                  80
Asp Cys Pro Lys Ile Thr Gln His Trp Arg Lys Trp Met Arg Arg Gly
                85                  90                  95


<210> 205
<211> 428
<212> DNA
<213> Homo sapiens

<400> 205
ctgggtctgt ctctgccacc tggtctgcca cagatccatg atgtgcagtt ctctggagca 60
ggcgctggct gtgctggtca ctaccttcca caagtactcc tgccaagagg gcgacaagtt 120
caagctgagt aaggggaaa tgaaggaact tctgcacaag gagctgccca gctttgtggg 180
ggagaaagtg atgaggagg ggctgaagaa gctgatgggc agcctggatg agaacagtga 240
ccagcaggtg gacttccagg agtatgctgt tttcctggca ctcatcactg tcatgtgcaa 300
tgacttcttc cagggctgcc cagaccgacc ctgaagcaga actcttgact tcctgccatg 360
gatctcttgg gcccaggact gttgatgcct ttgagttttg tattcaataa actttttttg 420
tctgttga                                                          428


<210> 206
<211> 97
<212> PRT
<213> Homo sapiens

<400> 206
Met Cys Ser Ser Leu Glu Gln Ala Leu Ala Val Leu Val Thr Thr Phe
```

```
 1                5                    10                    15
His Lys Tyr Ser Cys Gln Glu Gly Asp Lys Phe Lys Leu Ser Lys Gly
               20                    25                    30
Glu Met Lys Glu Leu Leu His Lys Glu Leu Pro Ser Phe Val Gly Glu
           35                    40                    45
Lys Val Asp Glu Glu Gly Leu Lys Lys Leu Met Gly Ser Leu Asp Glu
       50                    55                    60
Asn Ser Asp Gln Gln Val Asp Phe Gln Glu Tyr Ala Val Phe Leu Ala
65                    70                    75                    80
Leu Ile Thr Val Met Cys Asn Asp Phe Phe Gln Gly Cys Pro Asp Arg
               85                    90                    95
Pro
```

```
<210> 207
<211> 799
<212> DNA
<213> Homo sapiens

<400> 207
cactcccaaa gaactgggta ctcaacactg agcagatctg ttctttgagc taaaaaccat 60
gtgctgtacc aagagtttgc tcctggctgc tttgatgtca gtgctgctac tccacctctg 120
cggcgaatca gaagcagcaa gcaactttga ctgctgtctt ggatacacag accgtattct 180
tcatcctaaa tttattgtgg gcttcacacg gcagctggcc aatgaaggct gtgacatcaa 240
tgctatcatc tttcacacaa agaaaaagtt gtctgtgtgc gcaaatccaa aacagacttg 300
ggtgaaatat attgtgcgtc tcctcagtaa aaaagtcaag aacatgtaaa aactgtggct 360
tttctggaat ggaattggac atagcccaag aacagaaaga accttgctgg ggttggaggt 420
ttcacttgca catcatggag ggtttagtgc ttatctaatt tgtgcctcac tggacttgtc 480
caattaatga agttgattca tattgcatca tagtttgctt tgtttaagca tcacattaaa 540
gttaaactgt attttatgtt atttatagct gtaggttttc tgtgtttagc tatttaatac 600
taattttcca taagctattt tggtttagtg caaagtataa aattatattt ggggggggaat 660
aagattatat ggactttctt gcaagcaaca agctattttt taaaaaaact atttaacatt 720
cttttgttta tattgttttg tctcctaaat tgttgtaatt gcattataaa ataagaaaaa 780
cattaataag acaaatatt 799
```

```
<210> 208
<211> 96
<212> PRT
<213> Homo sapiens

<400> 208
Met Cys Cys Thr Lys Ser Leu Leu Leu Ala Ala Leu Met Ser Val Leu
1                5                    10                    15
Leu Leu His Leu Cys Gly Glu Ser Glu Ala Ala Ser Asn Phe Asp Cys
               20                    25                    30
Cys Leu Gly Tyr Thr Asp Arg Ile Leu His Pro Lys Phe Ile Val Gly
           35                    40                    45
Phe Thr Arg Gln Leu Ala Asn Glu Gly Cys Asp Ile Asn Ala Ile Ile
       50                    55                    60
Phe His Thr Lys Lys Lys Leu Ser Val Cys Ala Asn Pro Lys Gln Thr
65                    70                    75                    80
Trp Val Lys Tyr Ile Val Arg Leu Leu Ser Lys Lys Val Lys Asn Met
               85                    90                    95
```

```
<210> 209
<211> 2133
<212> DNA
```

<213> Homo sapiens

<400> 209
```
cgggagagcg cgctctgcct gccgcctgcc tgcctgccac tgagggttcc cagcaccatg 60
agggcctgga tcttctttct cctttgcctg gccgggaggg ccttggcagc ccctcagcaa 120
gaagccctgc ctgatgagac agaggtggtg gaagaaactg tggcagaggt gactgaggta 180
tctgtgggag ctaatcctgt ccaggtggaa gtaggagaat ttgatgatgg tgcagaggaa 240
accgaagagg aggtggtggc ggaaaatccc tgccagaacc accactgcaa acacggcaag 300
gtgtgcgagc tggatgagaa caacaccccc atgtgcgtgt gccaggaccc caccagctgc 360
ccagccccca ttggcgagtt tgagaaggtg tgcagcaatg acaacaagac cttcgactct 420
tcctgccact ctctttgccac aaagtgcacc ctggagggca ccaagaaggg ccacaagctc 480
cacctggact acatcgggcc ttgcaaatac atcccccctt gcctggactc tgagctgacc 540
gaattccccc tgcgcatgcg ggactggctc aagaacgtcc tggtcacccct gtatgagagg 600
gatgaggaca acaaccttct gactgagaag cagaagctgc gggtgaagaa gatccatgag 660
aatgagaagc gcctggaggc aggagaccac cccgtggagc tgctggcccg ggacttcgag 720
aagaactata acatgtacat cttccctgta cactggcagt cggccagct ggaccagcac 780
cccattgacg ggtacctctc ccacaccgag ctggctccac tgcgtgctcc cctcatcccc 840
atggagcatt gcaccacccg ctttttcgag acctgtgacc tggacaatga caagtacatc 900
gccctggatg agtgggccgg ctgcttcggc atcaagcaga aggatatcga caaggatctt 960
gtgatctaaa tccactcctt ccacagtacc ggattctctc tttaaccctc cccttcgtgt 1020
ttcccccaat gtttaaaatg tttggatggt ttgttgttct gcctggagac aaggtgctaa 1080
catagattta agtgaataca ttaacggtgc taaaaatgaa aattctaacc caagacatga 1140
cattcttagc tgtaacttaa ctattaaggc cttttccaca cgcattaata gtcccatttt 1200
tctcttgcca tttgtagctt tgcccattgt cttattggca catgggtgga cacggatctg 1260
ctgggctctg ccttaaacac acattgcagc ttcaactttt ctctttagtg ttctgtttga 1320
aactaatact taccgagtca gactttgtgt tcatttcatt tcagggtctt ggctgcctgt 1380
gggcttcccc aggtggcctg gaggtgggca aagggaagta acagacacac gatgttgtca 1440
aggatggttt tgggactaga ggctcagtgg tgggagagat ccctgcagaa tccaccaacc 1500
agaacgtggt ttgcctgagg ctgtaactga gagaaagatt ctggggctgt cttatgaaaa 1560
tatagacatt ctcacataag cccagttcat caccatttcc tcctttacct ttcagtgcag 1620
tttcttttca cattaggctg ttggttcaaa cttttgggag cacggactgt cagttctctg 1680
ggaagtggtc agcgcatcct gcagggcttc tcctcctctg tcttttggag aaccagggct 1740
cttctcaggg gctctaggga ctgccaggct gtttcagcca ggaaggccaa aatcaagagt 1800
gagatgtaga aagttgtaaa atagaaaaag tggagttggt gaatcggttg ttctttcctc 1860
acattggat gattgtcata aggttttag catgttcctc cttttcttca ccctcccctt 1920
tgttcttcta ttaatcaaga gaaacttcaa agttaatggg atggtcggat ctcacaggct 1980
gagaactcgt tcacctccaa gcatttcatg aaaaagctgc ttcttattaa tcatacaaac 2040
tctcaccatg atgtgaagag tttcacaaat ctttcaaaat aaaaagtaat gacttagaaa 2100
ctgaaaaaaa aaaaaaaaaa aaaaaaaaaa aaa 2133
```

<210> 210
<211> 303
<212> PRT
<213> Homo sapiens

<400> 210
```
Met Arg Ala Trp Ile Phe Phe Leu Leu Cys Leu Ala Gly Arg Ala Leu
 1               5                  10                  15
Ala Ala Pro Gln Gln Glu Ala Leu Pro Asp Glu Thr Glu Val Val Glu
            20                  25                  30
Glu Thr Val Ala Glu Val Thr Glu Val Ser Val Gly Ala Asn Pro Val
        35                  40                  45
Gln Val Glu Val Gly Glu Phe Asp Asp Gly Ala Glu Glu Thr Glu Glu
    50                  55                  60
Glu Val Val Ala Glu Asn Pro Cys Gln Asn His Cys Lys His Gly
65                  70          75                  80
Lys Val Cys Glu Leu Asp Glu Asn Asn Thr Pro Met Cys Val Cys Gln
                85                  90                  95
Asp Pro Thr Ser Cys Pro Ala Pro Ile Gly Glu Phe Glu Lys Val Cys
```

```
                100                    105                   110
Ser Asn Asp Asn Lys Thr Phe Asp Ser Ser Cys His Phe Phe Ala Thr
        115                    120                   125
Lys Cys Thr Leu Glu Gly Thr Lys Lys Gly His Lys Leu His Leu Asp
    130                    135                   140
Tyr Ile Gly Pro Cys Lys Tyr Ile Pro Pro Cys Leu Asp Ser Glu Leu
145                    150                   155                   160
Thr Glu Phe Pro Leu Arg Met Arg Asp Trp Leu Lys Asn Val Leu Val
                165                    170                   175
Thr Leu Tyr Glu Arg Asp Glu Asp Asn Asn Leu Leu Thr Glu Lys Gln
            180                    185                   190
Lys Leu Arg Val Lys Lys Ile His Glu Asn Glu Lys Arg Leu Glu Ala
        195                    200                   205   ·
Gly Asp His Pro Val Glu Leu Leu Ala Arg Asp Phe Glu Lys Asn Tyr
    210                    215                   220
Asn Met Tyr Ile Phe Pro Val His Trp Gln Phe Gly Gln Leu Asp Gln
225                    230                   235                   240
His Pro Ile Asp Gly Tyr Leu Ser His Thr Glu Leu Ala Pro Leu Arg
                245                    250                   255
Ala Pro Leu Ile Pro Met Glu His Cys Thr Thr Arg Phe Phe Glu Thr
            260                    265                   270
Cys Asp Leu Asp Asn Asp Lys Tyr Ile Ala Leu Asp Glu Trp Ala Gly
        275                    280                   285
Cys Phe Gly Ile Lys Gln Lys Asp Ile Asp Lys Asp Leu Val Ile
    290                    295                   300
```

```
<210> 211
<211> 2228
<212> DNA
<213> Homo sapiens

<400> 211
ggtacagtca tcacaagcct gttcggcggg actgtgatgg ccagagagat gacgatctta 60
ggatcggctg ttttgactct cctgttggcc ggctatttgg cacaacagta tttaccattg 120
cctactccta aagtgattgg tattgatctt ggcaccacct attgttctgt tggggtgttt 180
tttcctggca caggaaaagt aaaggtgatt ccagatgaaa atgggcatat cagcataccc 240
agcatggtgt cttttactga caatgatgta tatgtgggat atgaaagcgt agagctggca 300
gattcaaatc ctcaaaacac aatatatgat gccaaaagat tcataggcaa gattttttacc 360
gcagaagagt tggaggctga aattggcaga tacccattta aggttttaaa caaaaatgga 420
atggttgagt tttctgtgac aagtaatgag accatcacag tgtccccaga atatgttggc 480
tctcgactat tgttgaagtt aaaggaaatg gcagaggcat atcttggaat gccagttgcc 540
aatgctgtca tttctgtacc agcagaattt gatctaaaac agagaaattc aacaattgaa 600
gctgctaacc ttgcaggact gaagattttg agggtaataa atgaacccac agcagcagct 660
atggcctatg gtctccacaa ggctgacgtc ttccacgtct tggtgataga cttgggcgga 720
ggaactctag atgtgtcttt actgaataaa caaggaggga tgtttctaac ccgagcaatg 780
tctggaaaca ataaacttgg aggacaggac ttcaatcaga gattgcttca gtacttatat 840
aaacagatct atcaaacata tggcttcgtg ccctctagga agaggaaat ccacagattg 900
agacaagctg tggaaatggt caaattaaat ctgactcttc atcaatctgc tcagttgtca 960
gtattactaa cggtggagga gcaggacagg aaggaacctc acagtagtga cactgaactg 1020
ccaaaagaca aactttcctc agcagatgac catcgcgtga acagtgggtt tggacgtggc 1080
ctttctgata agaaaagtgg agaaagtcag gttttatttg aaacagaaat atcacggaaa 1140
ctctttgata cccttaatga agacctcttt cagaaaatac tggtacccat tcagcaagta 1200
ttgaaagaag gccacctgga aaagactgag attgatgagg tggtttttagt tggggggctcc 1260
actcgtattc ctcggatccg tcaagtcatt caagagttct ttggaaaaga tcccaacaca 1320
tctgtagacc ctgacctagc agtagtaacg ggagtggcta tccaagcagg gattgatgga 1380
ggctcttggc tctccaagt cagtgcttta gaaattccca ataagcattt acaaaaaacc 1440
aacttcaact gaattctgca gaaataatgg ttatttgtga acttgtctga tgatctcttc 1500
ccatttatca gattaccttt tccacaaaag aaagtctcta aaatatcaca gatttaccta 1560
```

```
gagggcaaca tttagataca ggaaaatttt acatagtgtt ttgtcttagg attagacgtg 1620
accagattga tcctgtttga ttttggagag atcctattct aacaaatact ctaaatgat 1680
aaaattgagg tacaactctc ttaaaagagt atggataact atattttctg gattctggag 1740
gttgataacc atatgcactt aacattatat tctataaaca ttaagtagtg ccagttatga 1800
gattcccagt tcttactaaa ttgtattagc aggagctggt aattacttgt attatcacat 1860
gtaactaata atttgaacta tacttgaagg accgtgttga tgtcaggtat ttacagtggt 1920
tggaagatag cagtattatt agcataagct gcatacgtaa tattcagtaa ctgccatatt 1980
atataacaaa tttacattca caaattcagt atcctgttaa gtgtcatatt cttgtaatct 2040
gcattctcca ggagttttat gtgtttaata gatgaattta ttttatttct aaaggtattc 2100
aaatgtttca gcaccatata atagaaatac ccaattatat ·tctagttcct ttatgtcctg 2160
tacatcattc tctgcttgga tttccattat tctgtttggt tagagaataa aattggtaat 2220
tgcatttg                                                             2228
```

<210> 212
<211> 471
<212> PRT
<213> Homo sapiens

<400> 212

```
Met Ala Arg Glu Met Thr Ile Leu Gly Ser Ala Val Leu Thr Leu Leu
 1               5                  10                  15
Leu Ala Gly Tyr Leu Ala Gln Gln Tyr Leu Pro Leu Pro Thr Pro Lys
                20                  25                  30
Val Ile Gly Ile Asp Leu Gly Thr Thr Tyr Cys Ser Val Gly Val Phe
            35                  40                  45
Phe Pro Gly Thr Gly Lys Val Lys Val Ile Pro Asp Glu Asn Gly His
    50                  55                  60
Ile Ser Ile Pro Ser Met Val Ser Phe Thr Asp Asn Asp Val Tyr Val
65                  70                  75                  80
Gly Tyr Glu Ser Val Glu Leu Ala Asp Ser Asn Pro Gln Asn Thr Ile
                85                  90                  95
Tyr Asp Ala Lys Arg Phe Ile Gly Lys Ile Phe Thr Ala Glu Glu Leu
                100                 105                 110
Glu Ala Glu Ile Gly Arg Tyr Pro Phe Lys Val Leu Asn Lys Asn Gly
            115                 120                 125
Met Val Glu Phe Ser Val Thr Ser Asn Glu Thr Ile Thr Val Ser Pro
    130                 135                 140
Glu Tyr Val Gly Ser Arg Leu Leu Leu Lys Leu Lys Glu Met Ala Glu
145                 150                 155                 160
Ala Tyr Leu Gly Met Pro Val Ala Asn Ala Val Ile Ser Val Pro Ala
                165                 170                 175
Glu Phe Asp Leu Lys Gln Arg Asn Ser Thr Ile Glu Ala Ala Asn Leu
                180                 185                 190
Ala Gly Leu Lys Ile Leu Arg Val Ile Asn Glu Pro Thr Ala Ala Ala
            195                 200                 205
Met Ala Tyr Gly Leu His Lys Ala Asp Val Phe His Val Leu Val Ile
    210                 215                 220
Asp Leu Gly Gly Gly Thr Leu Asp Val Ser Leu Leu Asn Lys Gln Gly
225                 230                 235                 240
Gly Met Phe Leu Thr Arg Ala Met Ser Gly Asn Asn Lys Leu Gly Gly
                245                 250                 255
Gln Asp Phe Asn Gln Arg Leu Leu Gln Tyr Leu Tyr Lys Gln Ile Tyr
                260                 265                 270
Gln Thr Tyr Gly Phe Val Pro Ser Arg Lys Glu Glu Ile His Arg Leu
            275                 280                 285
Arg Gln Ala Val Glu Met Val Lys Leu Asn Leu Thr Leu His Gln Ser
        290                 295                 300
Ala Gln Leu Ser Val Leu Leu Thr Val Glu Glu Gln Asp Arg Lys Glu
305                 310                 315                 320
```

274

```
Pro His Ser Ser Asp Thr Glu Leu Pro Lys Asp Lys Leu Ser Ser Ala
            325                 330                 335
Asp Asp His Arg Val Asn Ser Gly Phe Gly Arg Gly Leu Ser Asp Lys
        340                 345                 350
Lys Ser Gly Glu Ser Gln Val Leu Phe Glu Thr Glu Ile Ser Arg Lys
        355                 360                 365
Leu Phe Asp Thr Leu Asn Glu Asp Leu Phe Gln Lys Ile Leu Val Pro
    370                 375                 380
Ile Gln Gln Val Leu Lys Glu Gly His Leu Glu Lys Thr Glu Ile Asp
385                 390                 395                 400
Glu Val Val Leu Val Gly Gly Ser Thr Arg Ile Pro Arg Ile Arg Gln
            405                 410                 415
Val Ile Gln Glu Phe Phe Gly Lys Asp Pro Asn Thr Ser Val Asp Pro
        420                 425                 430
Asp Leu Ala Val Val Thr Gly Val Ala Ile Gln Ala Gly Ile Asp Gly
        435                 440                 445
Gly Ser Trp Pro Leu Gln Val Ser Ala Leu Glu Ile Pro Asn Lys His
    450                 455                 460
Leu Gln Lys Thr Asn Phe Asn
465                 470
```

```
<210> 213
<211> 1224
<212> DNA
<213> Homo sapiens

<400> 213
ggccgggaga gtagcagtgc cttggacccc agctctcctc cccctttctc tctaaggatg 60
gcccagaagg agaactccta cccctggccc tacggccgac agacggctcc atctggcctg 120
agcaccctgc cccagcgagt cctccggaaa gagcctgtca ccccatctgc acttgtcctc 180
atgagccgct ccaatgtcca gcccacagct gcccctggcc agaaggtgat ggagaatagc 240
agtgggacac ccgacatctt aacgcggcac ttcacaattg atgactttga gattgggcgt 300
cctctgggca aaggcaagtt tggaaacgtg tacttggctc gggagaagaa aagccatttc 360
atcgtggcgc tcaaggtcct cttcaagtcc cagatagaga aggagggcgt ggagcatcag 420
ctgcgcagag agatcgaaat ccaggcccac ctgcaccatc ccaacatcct gcgtctctac 480
aactattttt atgaccggag gaggatctac ttgattctag agtatgcccc ccgcggggag 540
ctctacaagg agctgcagaa gagctgcaca tttgacgagc agcgaacagc cacgatcatg 600
gaggagttgg cagatgctct aatgtactgc atgggaagaa ggtgattca cagagacata 660
aagccagaaa atctgctctt agggctcaag ggagagctga agattgctga cttcggctgg 720
tctgtgcatg cgccctccct gaggaggaag acaatgtgtg gcaccctgga ctacctgccc 780
ccagagatga ttgaggggcg catgcacaat gagaaggtgg atctgtggtg cattggagtg 840
ctttgctatg agctgctggt ggggaaccca ccctttgaga gtgcatcaca caacgagacc 900
tatcgccgca tcgtcaaggt ggacctaaag ttccccgctt ctgtgcccac gggagcccag 960
gacctcatct ccaaactgct caggcataac ccctcggaac ggctgcccct ggcccaggtc 1020
tcagcccacc cttgggtccg ggccaactct cggagggtgc tgcctccctc tgcccttcaa 1080
tctgtcgcct gatggtccct gtcattcact cgggtgcgtg tgtttgtatg tctgtgtatg 1140
tataggggaa agaagggatc cctaactgtt cccttatctg ttttctacct cctcctttgt 1200
ttaataaagg ctgaagcttt ttgt                                     1224
```

```
<210> 214
<211> 344
<212> PRT
<213> Homo sapiens

<400> 214
Met Ala Gln Lys Glu Asn Ser Tyr Pro Trp Pro Tyr Gly Arg Gln Thr
 1               5                  10                  15
Ala Pro Ser Gly Leu Ser Thr Leu Pro Gln Arg Val Leu Arg Lys Glu
```

```
              20                    25                    30
Pro Val Thr Pro Ser Ala Leu Val Leu Met Ser Arg Ser Asn Val Gln
          35                    40                    45
Pro Thr Ala Ala Pro Gly Gln Lys Val Met Glu Asn Ser Ser Gly Thr
      50                    55                    60
Pro Asp Ile Leu Thr Arg His Phe Thr Ile Asp Asp Phe Glu Ile Gly
65                    70                    75                    80
Arg Pro Leu Gly Lys Gly Lys Phe Gly Asn Val Tyr Leu Ala Arg Glu
                  85                    90                    95
Lys Lys Ser His Phe Ile Val Ala Leu Lys Val Leu Phe Lys Ser Gln
              100                   105                   110
Ile Glu Lys Glu Gly Val Glu His Gln Leu Arg Arg Glu Ile Glu Ile
          115                   120                   125
Gln Ala His Leu His His Pro Asn Ile Leu Arg Leu Tyr Asn Tyr Phe
      130                   135                   140
Tyr Asp Arg Arg Arg Ile Tyr Leu Ile Leu Glu Tyr Ala Pro Arg Gly
145                   150                   155                   160
Glu Leu Tyr Lys Glu Leu Gln Lys Ser Cys Thr Phe Asp Glu Gln Arg
                  165                   170                   175
Thr Ala Thr Ile Met Glu Glu Leu Ala Asp Ala Leu Met Tyr Cys His
              180                   185                   190
Gly Lys Lys Val Ile His Arg Asp Ile Lys Pro Glu Asn Leu Leu Leu
          195                   200                   205
Gly Leu Lys Gly Glu Leu Lys Ile Ala Asp Phe Gly Trp Ser Val His
      210                   215                   220
Ala Pro Ser Leu Arg Arg Lys Thr Met Cys Gly Thr Leu Asp Tyr Leu
225                   230                   235                   240
Pro Pro Glu Met Ile Glu Gly Arg Met His Asn Glu Lys Val Asp Leu
                  245                   250                   255
Trp Cys Ile Gly Val Leu Cys Tyr Glu Leu Leu Val Gly Asn Pro Pro
              260                   265                   270
Phe Glu Ser Ala Ser His Asn Glu Thr Tyr Arg Arg Ile Val Lys Val
          275                   280                   285
Asp Leu Lys Phe Pro Ala Ser Val Pro Thr Gly Ala Gln Asp Leu Ile
      290                   295                   300
Ser Lys Leu Leu Arg His Asn Pro Ser Glu Arg Leu Pro Leu Ala Gln
305                   310                   315                   320
Val Ser Ala His Pro Trp Val Arg Ala Asn Ser Arg Arg Val Leu Pro
                  325                   330                   335
Pro Ser Ala Leu Gln Ser Val Ala
              340
```

<210> 215
<211> 1421
<212> DNA
<213> Homo sapiens

<400> 215
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acttactgcg ggacggcctt ggagagtact cgggttcgtg aacttcccgg aggcgcaatg 60
agctgcatta acctgcccac tgtgctgccc ggctccccca gcaagacccg ggggcagatc 120
caggtgattc tcggccgat gttctcagga aaaagcacag agttgatgag acgcgtccgt 180
cgcttccaga ttgctcagta caagtgcctg gtgatcaagt atgccaaaga cactcgctac 240
agcagcagct tctgcacaca tgaccggaac accatggagg cgctgcccgc ctgcctgctc 300
cgagacgtgg cccaggaggc cctgggcgtg ctgtcatag gcatcgacga ggggcagttt 360
ttccctgaca tcatggagtt ctgcgaggcc atggccaacg ccgggaagac cgtaattgtg 420
gctgcactgg atgggacctt ccagaggaag ccatttgggg ccatcctgaa cctggtgccg 480
ctggccgaga gcgtggtgaa gctgacggcg gtgtgcatgg agtgcttccg ggaagccgcc 540
tataccaaga ggctcggcac agagaaggag gtcgaggtga ttgggggagc agacaagtac 600
```

```
cactccgtgt gtcggctctg ctacttcaag aaggcctcag gccagcctgc cgggccggac 660
aacaaagaga actgcccagt gccaggaaag ccaggggaag ccgtggctgc caggaagctc 720
tttgccccac agcagattct gcaatgcagc cctgccaact gagggacctg caagggccgc 780
ccgctccctt cctgccactg ccgcctactg gacgctgccc tgcatgctgc ccagccactc 840
caggaggaag tcgggaggcg tggagggtga ccacaccttg gccttctggg aactctcctt 900
tgtgtggctg ccccacctgc cgcatgctcc ctcctctcct acccactggt ctgcttaaag 960
cttccctctc agctgctggg acgatcgccc aggctggagc tggccccgct tggtggcctg 1020
ggatctggca cactccctct ccttggggtg agggacagag ccccacgctg ttgacatcag 1080
cctgcttctt cccctctgcg gctttcactg ctgagtttct gttctccctg ggaagcctgt 1140
gccagcacct ttgagccttg ccccacactg aggcttaggc ctctctgcct gggatgggct 1200
cccaccctcc cctgaggatg gcctggattc acgccctctt gtttcctttt gggctcaaag 1260
cccttcctac ctctggtgat ggtttccaca ggaacaacag catctttcac caagatgggt 1320
ggcaccaacc ttgctgggac ttggatccca ggggcttatc tcttcaagtg tggagagggc 1380
agggtccacg cctctgctgt agcttatgaa attaactaat t              1421
```

<210> 216
<211> 234
<212> PRT
<213> Homo sapiens

<400> 216

```
Met Ser Cys Ile Asn Leu Pro Thr Val Leu Pro Gly Ser Pro Ser Lys
1               5                   10                  15
Thr Arg Gly Gln Ile Gln Val Ile Leu Gly Pro Met Phe Ser Gly Lys
            20                  25                  30
Ser Thr Glu Leu Met Arg Arg Val Arg Arg Phe Gln Ile Ala Gln Tyr
        35                  40                  45
Lys Cys Leu Val Ile Lys Tyr Ala Lys Asp Thr Arg Tyr Ser Ser Ser
    50                  55                  60
Phe Cys Thr His Asp Arg Asn Thr Met Glu Ala Leu Pro Ala Cys Leu
65                  70                  75                  80
Leu Arg Asp Val Ala Gln Glu Ala Leu Gly Val Ala Val Ile Gly Ile
                85                  90                  95
Asp Glu Gly Gln Phe Phe Pro Asp Ile Met Glu Phe Cys Glu Ala Met
            100                 105                 110
Ala Asn Ala Gly Lys Thr Val Ile Val Ala Ala Leu Asp Gly Thr Phe
        115                 120                 125
Gln Arg Lys Pro Phe Gly Ala Ile Leu Asn Leu Val Pro Leu Ala Glu
    130                 135                 140
Ser Val Val Lys Leu Thr Ala Val Cys Met Glu Cys Phe Arg Glu Ala
145                 150                 155                 160
Ala Tyr Thr Lys Arg Leu Gly Thr Glu Lys Glu Val Glu Val Ile Gly
                165                 170                 175
Gly Ala Asp Lys Tyr His Ser Val Cys Arg Leu Cys Tyr Phe Lys Lys
            180                 185                 190
Ala Ser Gly Gln Pro Ala Gly Pro Asp Asn Lys Glu Asn Cys Pro Val
        195                 200                 205
Pro Gly Lys Pro Gly Glu Ala Val Ala Ala Arg Lys Leu Phe Ala Pro
    210                 215                 220
Gln Gln Ile Leu Gln Cys Ser Pro Ala Asn
225                 230
```

<210> 217
<211> 2307
<212> DNA
<213> Homo sapiens

<220>

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<221> misc_feature
<222> 1691, 1698, 1705, 1708, 1709, 1713, 1717, 1720, 1724, 1728,
1733, 1741, 1746, 1748, 1755, 1770, 1774, 1791, 1802, 1821,
1838, 1856, 1859, 1864, 1908, 1959, 1997, 2012, 2038, 2143
<223> n = A,T,C or G

<400> 217
agtcgacccc gcgtccggtt ttaatcaagc tgcccaaagt cccccaatca ctcctggaat 60
acacagagag aggcagcagc ttgctcagcg gacaaggatg ctgggcgtga gggaccaagg 120
cctgccctgc actcgggcct cctccagcca gtgctgacca gggacttctg acctgctggc 180
cagccaggac ctgtgtgggg aggccctcct gctgccttgg ggtgacaatc tcagctccag 240
gctacaggga gaccgggagg atcacagagc cagcatgtta caggatcctg acagtgatca 300
acctctgaac agcctcgatg tcaaacccct gcgcaaaccc cgtatcccca tggagacctt 360
cagaaaggtg gggatcccca tcatcatagc actactgagc ctggcgagta tcatcattgt 420
ggttgtcctc atcaaggtga ttctggataa atactacttc ctctgcgggc agcctctcca 480
cttcatcccg aggaagcagc tgtgtgacgg agagctggac tgtcccttgg gggaggacga 540
ggagcactgt gtcaagagct cccccgaagg gcctgcagtg gcagtccgcc tctccaagga 600
ccgatccaca ctgcaggtgc tggactcggc cacagggaac tggttctctg cctgtttcga 660
caacttcaca gaagctctcg ctgagacagc ctgtaggcag atgggctaca gcagcaaacc 720
cactttcaga gctgtggaga ttggcccaga ccaggatctg gatgttgttg aaatcacaga 780
aaacagccag gagcttcgca tgcggaactc aagtgggccc tgtctctcag gctccctggt 840
ctccctgcac tgtcttgcct gtgggaagag cctgaagacc ccccgtgtgg tgggtgggga 900
ggaggcctct gtggattctt ggccttggca ggtcagcatc cagtacgaca aacagcacgt 960
ctgtggaggg agcatcctgg accccactg ggtcctcacg gcagcccact gcttcaggaa 1020
acataccgat gtgttcaact ggaaggtgcg ggcaggctca gacaaactgg gcagcttccc 1080
atccctggct gtggccaaga tcatcatcat tgaattcaac cccatgtacc ccaaagacaa 1140
tgacatcgcc ctcatgaagc tgcagttccc actcactttc tcaggcacag tcaggcccat 1200
ctgtctgccc ttctttgatg aggagctcac tccagccacc ccactctgga tcattggatg 1260
gggctttacg aagcagaatg gagggaagat gtctgacata ctgctgcagg cgtcagtcca 1320
ggtcattgac agcacacggt gcaatgcaga cgatgcgtac cagggggaag tcaccgagaa 1380
gatgatgtgt gcaggcatcc cggaaggggg tgtggacacc tgccagggtg acagtggtgg 1440
gcccctgatg taccaatctg accagtggca tgtggtgggc atcgttagct ggggctatgg 1500
ctgcgggggc ccgagcaccc caggagtata caccaaggtc tcagcctatc tcaactggat 1560
ctacaatgtc tggaaggctg agctgtaatg ctgctgcccc tttgcagtgc tgggagccgc 1620
ttccttcctg ccctgcccac ctggggatcc cccaaagtca gacacagagc aagagtcccc 1680
ttgggtacac nccctctngc ccacnagnnc ctncagncan tttncttngg agncagcaaa 1740
ngggcncntc aattncctgt aagagacccn tcgncagccc agaggcgccc nagaggaagt 1800
cnagcagccc tagctcggcc nacacttggt gctcccangc atcccaggga gagacnacna 1860
gccncactga acaaggtctc aggggtattg ctaagccaag aaggaacntt tcccacacta 1920
ctgaatggaa gcaggctgtc ttgtaaaagc ccagatcanc tgtgggctgg agaggagaag 1980
gaaagggtct gcgccangcc ctgtccgtct tncacccatc cccaagccta ctagagcnaa 2040
gaaaccagtt gtaatataaa atgcactgcc ctactgttgg tatgactacc gttacctact 2100
gttgtcattg ttattacagc tatggccact attattaaag agnctgtgta acatcaaaaa 2160
aaaaaaaaaa aaaaaaaaaa araaaaaaa ataaataaaa aaaaactcga gggggggccc 2220
ggtacccaat tcgccctata gtgagtcgta ttacaattca ctggccgtcg ttttacaacg 2280
tcgtgactgg gaaaaccctg gcgttac                                     2307

<210> 218
<211> 428
<212> PRT
<213> Homo sapiens

<400> 218
Met Leu Gln Asp Pro Asp Ser Asp Gln Pro Leu Asn Ser Leu Asp Val
1               5                   10                  15
Lys Pro Leu Arg Lys Pro Arg Ile Pro Met Glu Thr Phe Arg Lys Val
            20                  25                  30
Gly Ile Pro Ile Ile Ile Ala Leu Leu Ser Leu Ala Ser Ile Ile Ile
            35                  40                  45
```

```
Val Val Val Leu Ile Lys Val Ile Leu Asp Lys Tyr Tyr Phe Leu Cys
    50              55              60
Gly Gln Pro Leu His Phe Ile Pro Arg Lys Gln Leu Cys Asp Gly Glu
65              70              75              80
Leu Asp Cys Pro Leu Gly Glu Asp Glu Glu His Cys Val Lys Ser Phe
            85              90              95
Pro Glu Gly Pro Ala Val Ala Val Arg Leu Ser Lys Asp Arg Ser Thr
            100             105             110
Leu Gln Val Leu Asp Ser Ala Thr Gly Asn Trp Phe Ser Ala Cys Phe
        115             120             125
Asp Asn Phe Thr Glu Ala Leu Ala Glu Thr Ala Cys Arg Gln Met Gly
    130             135             140
Tyr Ser Ser Lys Pro Thr Phe Arg Ala Val Glu Ile Gly Pro Asp Gln
145             150             155             160
Asp Leu Asp Val Val Glu Ile Thr Glu Asn Ser Gln Glu Leu Arg Met
            165             170             175
Arg Asn Ser Ser Gly Pro Cys Leu Ser Gly Ser Leu Val Ser Leu His
            180             185             190
Cys Leu Ala Cys Gly Lys Ser Leu Lys Thr Pro Arg Val Val Gly Gly
        195             200             205
Glu Glu Ala Ser Val Asp Ser Trp Pro Trp Gln Val Ser Ile Gln Tyr
    210             215             220
Asp Lys Gln His Val Cys Gly Gly Ser Ile Leu Asp Pro His Trp Val
225             230             235             240
Leu Thr Ala Ala His Cys Phe Arg Lys His Thr Asp Val Phe Asn Trp
            245             250             255
Lys Val Arg Ala Gly Ser Asp Lys Leu Gly Ser Phe Pro Ser Leu Ala
            260             265             270
Val Ala Lys Ile Ile Ile Ile Glu Phe Asn Pro Met Tyr Pro Lys Asp
        275             280             285
Asn Asp Ile Ala Leu Met Lys Leu Gln Phe Pro Leu Thr Phe Ser Gly
    290             295             300
Thr Val Arg Pro Ile Cys Leu Pro Phe Phe Asp Glu Glu Leu Thr Pro
305             310             315             320
Ala Thr Pro Leu Trp Ile Ile Gly Trp Gly Phe Thr Lys Gln Asn Gly
            325             330             335
Gly Lys Met Ser Asp Ile Leu Leu Gln Ala Ser Val Gln Val Ile Asp
            340             345             350
Ser Thr Arg Cys Asn Ala Asp Asp Ala Tyr Gln Gly Glu Val Thr Glu
        355             360             365
Lys Met Met Cys Ala Gly Ile Pro Glu Gly Gly Val Asp Thr Cys Gln
370             375             380
Gly Asp Ser Gly Gly Pro Leu Met Tyr Gln Ser Asp Gln Trp His Val
385             390             395             400
Val Gly Ile Val Ser Trp Gly Tyr Gly Cys Gly Gly Pro Ser Thr Pro
            405             410             415
Gly Val Tyr Thr Lys Val Ser Ala Tyr Leu Asn Trp
            420             425
```

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<210> 219
<211> 556
<212> DNA
<213> Homo sapiens

<400> 219
acaactcggt ggtggccact gcgcagacca gacttcgctc gtactcgtgc gcctcgcttc 60
gcttttcctc cgcaaccatg tctgacaaac ccgatatggc tgagatcgag aaattcgata 120
agtcgaaact gaagaagaca gagacgcaag agaaaaatcc actgccttcc aaagaaacga 180
```

```
ttgaacagga gaagcaagca ggcgaatcgt aatgaggcgt gcgccgccaa tatgcactgt 240
acattccaca agcattgcct tcttatttta cttcttttag ctgtttaact ttgtaagatg 300
caaagaggtt ggatcaagtt taaatgactg tgctgcccct ttcacatcaa agaactactg 360
acaacgaagg ccgcgctgcc tttcccatct gtctatctat ctggctggca gggaaggaaa 420
gaacttgcat gttggtgaag gaagaagtgg ggtggaagaa gtggggtggg acgacagtga 480
aatctagagt aaaaccaagc tggcccaagt gtcctgcagg ctgtaatgca gtttaatcag 540
agtgccattt tttttt                                                  556
```

<210> 220
<211> 44
<212> PRT
<213> Homo sapiens

<400> 220

```
Met Ser Asp Lys Pro Asp Met Ala Glu Ile Glu Lys Phe Asp Lys Ser
1               5                   10                  15
Lys Leu Lys Lys Thr Glu Thr Gln Glu Lys Asn Pro Leu Pro Ser Lys
            20                  25                  30
Glu Thr Ile Glu Gln Glu Lys Gln Ala Gly Glu Ser
        35                  40
```

<210> 221
<211> 4792
<212> DNA
<213> Homo sapiens

<400> 221

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ggaccaccca gtaccgatcc cttcacgacc gtcaccatgg aagtgtcacc attgcagcct 60
gtaaatgaaa atatgcaagt caacaaaata aagaaaatg aagatgctaa gaaaagactg 120
tctgttgaaa gaatctatca aaagaaaaca caattggaac atattttgct ccgcccagac 180
acctacattg gttctgtgga attagtgacc cagcaaatgt gggtttacga tgaagatgtt 240
ggcattaact atagggaagt cacttttgtt cctggtttgt acaaaatctt tgatgagatt 300
ctagttaatg ctgcggacaa caaacaaagg gacccaaaaa tgtcttgtat tagagtcaca 360
attgatccgg aaaacaattt aattagtata tggaataatg aaaaggtat tcctgttgtt 420
gaacacaaag ttgaaaagat gtatgtccca gctctcatat ttggacagct cctaacttct 480
agtaactatg atgatgatga aaagaaagtg acaggtggtc gaaatggcta tggagccaaa 540
ttgtgtaaca tattcagtac caaatttact gtggaaacag ccagtagaga atacaagaaa 600
atgttcaaac agacatggat ggataatatg ggaagagctg gtgagatgga actcaagccc 660
ttcaatggag aagattatac atgtatcacc tttcagcctg atttgtctaa gtttaaaatg 720
caaagcctgg acaaagatat tgttgcacta atggtcagaa gagcatatga tattgctgga 780
tccaccaaag atgtcaaagt ctttcttaat ggaaataaac tgccagtaaa aggatttcgt 840
agttatgtgg acatgtattt gaaggacaag ttggatgaaa ctggtaactc cttgaaagta 900
atacatgaac aagtaaacca caggtgggaa gtgtgtttaa ctatgagtga aaaaggcttt 960
cagcaaatta gctttgtcaa cagcattgct acatccaagg gtggcagaca tgttgattat 1020
gtagctgatc agattggtac taaacttgtt gatgttgtga agaagaagaa caagggtggt 1080
gttgcagtaa aagcacatca ggtgaaaaat cacatgtgga tttttgtaaa tgccttaatt 1140
gaaaacccaa ccttttgactc tcagacaaaa gaaaacatga ctttacaacc caagagcttt 1200
ggatcaacat gccaattgag tgaaaaattt atcaaagctg ccattggctg tggtattgta 1260
gaaagcatac taaactgggt gaagtttaag gcccaagtcc agttaaacaa gaagtgttca 1320
gctgtaaaac ataatagaat caagggaatt cccaaactcg atgatgccaa tgatgcaggg 1380
ggccgaaact ccactgagtg tacgcttatc ctgactgagg gagattcagc caaaactttg 1440
gctgtttcag gccttggtgt ggttgggaga gacaaatatg gggtttttccc tcttagagga 1500
aaaatactca atgttcgaga gcttctcat aagcagatca tggaaaatgc tgagattaac 1560
aatatcatca gagattgtggg tcttcagtac aagaaaaact atgaagatga agattcattg 1620
aagacgcttc gttatgggaa gataatgatt atgacagatc aggaccaaga tggttcccac 1680
atcaaaggct gctgattaa ttttatccat cacaactggc cctctcttct gcgacatcgt 1740
tttctggagg aatttatcac tcccattgta aaggtatcta aaaacaagca agaaatggca 1800
ttttacagcc ttcctgaatt tgaagagtgg aagagttcta ctccaaatca taaaaaatgg 1860
```

280

```
aaagtcaaat attacaaagg tttgggcacc agcacatcaa aggaagctaa agaatacttt 1920
gcagatatga aaagacatcg tatccagttc aaatattctg gtcctgaaga tgatgctgct 1980
atcagcctgg cctttagcaa aaaacagata gatgatcgaa aggaatggtt aactaatttc 2040
atggaggata gaagacaacg aaagttactt gggcttcctg aggattactt gtatggacaa 2100
actaccacat atctgacata taatgacttc atcaacaagg aacttatctt gttctcaaat 2160
tctgataacg agagatctat cccttctatg gtggatggtt tgaaaccagg tcagagaaag 2220
gttttgttta cttgcttcaa acggaatgac aagcgagaag taaaggttgc ccaattagct 2280
ggatcagtgg ctgaaatgtc ttcttatcat catggtgaga tgtcactaat gatgaccatt 2340
atcaatttgg ctcagaattt tgtgggtagc aataatctaa acctcttgca gcccattggt 2400
cagtttggta ccaggctaca tggtggcaag gattctgcta gtccacgata catctttaca 2460
atgctcagct ctttggctcg attgttattt ccaccaaaag atgatcacac gttgaagttt 2520
ttatatgatg acaaccagcg tgttgagcct gaatggtaca ttcctattat tcccatggtg 2580
ctgataaatg gtgctgaagg aatcggtact gggtggtcct gcaaaatccc caactttgat 2640
gtgcgtgaaa ttgtaaataa catcaggcgt ttgatggatg gagaagaacc tttgccaatg 2700
cttccaagtt acaagaactt caagggtact attgaagaac tggctccaaa tcaatatgtg 2760
attagtggtg aagtagctat tcttaattct acaaccattg aaatctcaga gcttcccgtc 2820
agaacatgga cccagacata caaagaacaa gttctagaac ccatgttgaa tggcaccgag 2880
aagacacctc ctctcataac agactatagg gaataccata cagataccac tgtgaaattt 2940
gttgtgaaga tgactgaaga aaaactggca gaggcagaga gagttggact acacaaagtc 3000
ttcaaactcc aaactagtct cacatgcaac tctatggtgc ttttttgacca cgtaggctgt 3060
ttaaagaaat atgacacggt gttggatatt ctaagagact ttttttgaact cagacttaaa 3120
tattatggat taagaaaaga atggctccta ggaatgcttg gtgctgaatc tgctaaactg 3180
aataatcagg ctcgctttat cttagagaaa atagatggca aaataatcat tgaaaataag 3240
cctaagaaag aattaattaa agttctgatt cagaggggat atgattcgga tcctgtgaag 3300
gcctggaaag aagcccagca aaaggttcca gatgaagaag aaaatgaaga gagtgacaac 3360
gaaaaggaaa ctgaaaagag tgactccgta acagattctg gaccaacctt caactatctt 3420
cttgatatgc cccttggta tttaaccaag gaaaagaaag atgaactctg caggctaaga 3480
aatgaaaaag aacaagagct ggacacatta aaaagaaaga gtccatcaga tttgtggaaa 3540
gaagacttgg ctacatttat tgaagaattg gaggctgttg aagccaagga aaaacaagat 3600
gaacaagtcg gacttcctgg gaaaggggggg aaggccaagg ggaaaaaaac acaaatggct 3660
gaagttttgc cttctccgcg tggtcaaaga gtcattccac gaataaccat agaaatgaaa 3720
gcagaggcag aaaagaaaaa taaaaagaaa attaagaatg aaaatactga aggaagccct 3780
caagaagatg gtgtggaact agaaggccta aaacaaagat tagaaaagaa acagaaaaga 3840
gaaccaggta caaagacaaa gaaacaaact acattggcat ttaagccaat caaaaaagga 3900
aagaagagaa atccctggcc tgattcagaa tcagatagga gcagtgacga aagtaatttt 3960
gatgtccctc cacgagaaac agagccacgg agagcagcaa caaaaacaaa attcacaatg 4020
gatttggatt cagatgaaga tttctcagat tttgatgaaa aaactgatga tgaagatttt 4080
gtcccatcag atgctagtcc acctaagacc aaaacttccc caaaacttag taacaaagaa 4140
ctgaaccac agaaaagtgt cgtgtcagac cttgaagctg atgatgttaa gggcagtgta 4200
ccactgtctt caagccctcc tgctacacat ttcccagatg aaactgaaat tacaaaccca 4260
gttcctaaaa agaatgtgac agtgaagaag acagcagcaa aaagtcagtc ttccacctcc 4320
actaccggtg ccaaaaaaag ggctgcccca aaaġgaacta aaagggatcc agctttgaat 4380
tctggtgtct ctcaaaagcc tgatcctgcc aaaaccaaga atcgccgcaa aaggaagcca 4440
tccacttctg atgattctga ctctaatttt gagaaaattg tttcgaaagc agtcacaagc 4500
aagaaatcca aggggggagag tgatgacttc catatggact ttgactcagc tgtggctcct 4560
cgggcaaaat ctgtacgggc aaagaaacct ataaagtacc tggaagagtc agatgaagat 4620
gatctgtttt aaaatgtgag gcgattattt taagtaatta tcttaccaag cccaagactg 4680
gtttttaaagt tacctgaagc tcttaacttc ctcccctctg aatttagttt ggggaaggtg 4740
tttttagtac aagacatcaa agtgaagtaa agcccaagtg ttctttagct tt        4792
```

<210> 222
<211> 1531
<212> PRT
<213> Homo sapiens

<400> 222
Met Glu Val Ser Pro Leu Gln Pro Val Asn Glu Asn Met Gln Val Asn
1   5   10   15
Lys Ile Lys Lys Asn Glu Asp Ala Lys Lys Arg Leu Ser Val Glu Arg

281

```
                  20                    25                    30
Ile Tyr Gln Lys Lys Thr Gln Leu Glu His Ile Leu Leu Arg Pro Asp
             35                    40                    45
Thr Tyr Ile Gly Ser Val Glu Leu Val Thr Gln Gln Met Trp Val Tyr
         50                    55                    60
Asp Glu Asp Val Gly Ile Asn Tyr Arg Glu Val Thr Phe Val Pro Gly
65                    70                    75                    80
Leu Tyr Lys Ile Phe Asp Glu Ile Leu Val Asn Ala Ala Asp Asn Lys
                 85                    90                    95
Gln Arg Asp Pro Lys Met Ser Cys Ile Arg Val Thr Ile Asp Pro Glu
             100                   105                   110
Asn Asn Leu Ile Ser Ile Trp Asn Asn Gly Lys Gly Ile Pro Val Val
         115                   120                   125
Glu His Lys Val Glu Lys Met Tyr Val Pro Ala Leu Ile Phe Gly Gln
     130                   135                   140
Leu Leu Thr Ser Ser Asn Tyr Asp Asp Asp Glu Lys Lys Val Thr Gly
145                   150                   155                   160
Gly Arg Asn Gly Tyr Gly Ala Lys Leu Cys Asn Ile Phe Ser Thr Lys
                 165                   170                   175
Phe Thr Val Glu Thr Ala Ser Arg Glu Tyr Lys Lys Met Phe Lys Gln
             180                   185                   190
Thr Trp Met Asp Asn Met Gly Arg Ala Gly Glu Met Glu Leu Lys Pro
             195                   200                   205
Phe Asn Gly Glu Asp Tyr Thr Cys Ile Thr Phe Gln Pro Asp Leu Ser
     210                   215                   220
Lys Phe Lys Met Gln Ser Leu Asp Lys Asp Ile Val Ala Leu Met Val
225                   230                   235                   240
Arg Arg Ala Tyr Asp Ile Ala Gly Ser Thr Lys Asp Val Lys Val Phe
             245                   250                   255
Leu Asn Gly Asn Lys Leu Pro Val Lys Gly Phe Arg Ser Tyr Val Asp
         260                   265                   270
Met Tyr Leu Lys Asp Lys Leu Asp Glu Thr Gly Asn Ser Leu Lys Val
     275                   280                   285
Ile His Glu Gln Val Asn His Arg Trp Glu Val Cys Leu Thr Met Ser
     290                   295                   300
Glu Lys Gly Phe Gln Gln Ile Ser Phe Val Asn Ser Ile Ala Thr Ser
305                   310                   315                   320
Lys Gly Gly Arg His Val Asp Tyr Val Ala Asp Gln Ile Val Thr Lys
             325                   330                   335
Leu Val Asp Val Val Lys Lys Lys Asn Lys Gly Gly Val Ala Val Lys
             340                   345                   350
Ala His Gln Val Lys Asn His Met Trp Ile Phe Val Asn Ala Leu Ile
     355                   360                   365
Glu Asn Pro Thr Phe Asp Ser Gln Thr Lys Glu Asn Met Thr Leu Gln
     370                   375                   380
Pro Lys Ser Phe Gly Ser Thr Cys Gln Leu Ser Glu Lys Phe Ile Lys
385                   390                   395                   400
Ala Ala Ile Gly Cys Gly Ile Val Glu Ser Ile Leu Asn Trp Val Lys
             405                   410                   415
Phe Lys Ala Gln Val Gln Leu Asn Lys Lys Cys Ser Ala Val Lys His
             420                   425                   430
Asn Arg Ile Lys Gly Ile Pro Lys Leu Asp Asp Ala Asn Asp Ala Gly
             435                   440                   445
Gly Arg Asn Ser Thr Glu Cys Thr Leu Ile Leu Thr Glu Gly Asp Ser
     450                   455                   460
Ala Lys Thr Leu Ala Val Ser Gly Leu Gly Val Gly Arg Asp Lys
465                   470                   475                   480
Tyr Gly Val Phe Pro Leu Arg Gly Lys Ile Leu Asn Val Arg Glu Ala
             485                   490                   495
```

```
Ser His Lys Gln Ile Met Glu Asn Ala Glu Ile Asn Asn Ile Ile Lys
            500             505             510
Ile Val Gly Leu Gln Tyr Lys Lys Asn Tyr Glu Asp Glu Asp Ser Leu
            515             520             525
Lys Thr Leu Arg Tyr Gly Lys Ile Met Ile Met Thr Asp Gln Asp Gln
    530             535             540
Asp Gly Ser His Ile Lys Gly Leu Leu Ile Asn Phe Ile His His Asn
545             550             555             560
Trp Pro Ser Leu Leu Arg His Arg Phe Leu Glu Glu Phe Ile Thr Pro
            565             570             575
Ile Val Lys Val Ser Lys Asn Lys Gln Glu Met Ala Phe Tyr Ser Leu
            580             585             590
Pro Glu Phe Glu Glu Trp Lys Ser Ser Thr Pro Asn His Lys Lys Trp
       595             600             605
Lys Val Lys Tyr Tyr Lys Gly Leu Gly Thr Ser Thr Ser Lys Glu Ala
    610             615             620
Lys Glu Tyr Phe Ala Asp Met Lys Arg His Arg Ile Gln Phe Lys Tyr
625             630             635             640
Ser Gly Pro Glu Asp Asp Ala Ala Ile Ser Leu Ala Phe Ser Lys Lys
            645             650             655
Gln Ile Asp Asp Arg Lys Glu Trp Leu Thr Asn Phe Met Glu Asp Arg
            660             665             670
Arg Gln Arg Lys Leu Leu Gly Leu Pro Glu Asp Tyr Leu Tyr Gly Gln
    675             680             685
Thr Thr Thr Tyr Leu Thr Tyr Asn Asp Phe Ile Asn Lys Glu Leu Ile
    690             695             700
Leu Phe Ser Asn Ser Asp Asn Glu Arg Ser Ile Pro Ser Met Val Asp
705             710             715             720
Gly Leu Lys Pro Gly Gln Arg Lys Val Leu Phe Thr Cys Phe Lys Arg
            725             730             735
Asn Asp Lys Arg Glu Val Lys Val Ala Gln Leu Ala Gly Ser Val Ala
            740             745             750
Glu Met Ser Ser Tyr His His Gly Glu Met Ser Leu Met Met Thr Ile
            755             760             765
Ile Asn Leu Ala Gln Asn Phe Val Gly Ser Asn Asn Leu Asn Leu Leu
    770             775             780
Gln Pro Ile Gly Gln Phe Gly Thr Arg Leu His Gly Gly Lys Asp Ser
785             790             795             800
Ala Ser Pro Arg Tyr Ile Phe Thr Met Leu Ser Ser Leu Ala Arg Leu
            805             810             815
Leu Phe Pro Pro Lys Asp Asp His Thr Leu Lys Phe Leu Tyr Asp Asp
            820             825             830
Asn Gln Arg Val Glu Pro Glu Trp Tyr Ile Pro Ile Ile Pro Met Val
            835             840             845
Leu Ile Asn Gly Ala Glu Gly Ile Gly Thr Gly Trp Ser Cys Lys Ile
    850             855             860
Pro Asn Phe Asp Val Arg Glu Ile Val Asn Asn Ile Arg Arg Leu Met
865             870             875             880
Asp Gly Glu Glu Pro Leu Pro Met Leu Pro Ser Tyr Lys Asn Phe Lys
            885             890             895
Gly Thr Ile Glu Glu Leu Ala Pro Asn Gln Tyr Val Ile Ser Gly Glu
            900             905             910
Val Ala Ile Leu Asn Ser Thr Thr Ile Glu Ile Ser Glu Leu Pro Val
    915             920             925
Arg Thr Trp Thr Gln Thr Tyr Lys Glu Gln Val Leu Glu Pro Met Leu
    930             935             940
Asn Gly Thr Glu Lys Thr Pro Pro Leu Ile Thr Asp Tyr Arg Glu Tyr
945             950             955             960
His Thr Asp Thr Thr Val Lys Phe Val Val Lys Met Thr Glu Glu Lys
```

```
              965                   970                   975
Leu Ala Glu Ala Glu Arg Val Gly Leu His Lys Val Phe Lys Leu Gln
              980                   985                   990
Thr Ser Leu Thr Cys Asn Ser Met Val Leu Phe Asp His Val Gly Cys
              995                  1000                  1005
Leu Lys Lys Tyr Asp Thr Val Leu Asp Ile Leu Arg Asp Phe Phe Glu
             1010                  1015                  1020
Leu Arg Leu Lys Tyr Tyr Gly Leu Arg Lys Glu Trp Leu Leu Gly Met
1025                   1030                  1035                  1040
Leu Gly Ala Glu Ser Ala Lys Leu Asn Asn Gln Ala Arg Phe Ile Leu
                 1045                  1050                  1055
Glu Lys Ile Asp Gly Lys Ile Ile Ile Glu Asn Lys Pro Lys Lys Glu
                 1060                  1065                  1070
Leu Ile Lys Val Leu Ile Gln Arg Gly Tyr Asp Ser Asp Pro Val Lys
             1075                  1080                  1085
Ala Trp Lys Glu Ala Gln Gln Lys Val Pro Asp Glu Glu Glu Asn Glu
             1090                  1095                  1100
Glu Ser Asp Asn Glu Lys Glu Thr Glu Lys Ser Asp Ser Val Thr Asp
1105                   1110                  1115                  1120
Ser Gly Pro Thr Phe Asn Tyr Leu Leu Asp Met Pro Leu Trp Tyr Leu
                 1125                  1130                  1135
Thr Lys Glu Lys Lys Asp Glu Leu Cys Arg Leu Arg Asn Glu Lys Glu
                 1140                  1145                  1150
Gln Glu Leu Asp Thr Leu Lys Arg Lys Ser Pro Ser Asp Leu Trp Lys
             1155                  1160                  1165
Glu Asp Leu Ala Thr Phe Ile Glu Glu Leu Glu Ala Val Glu Ala Lys
1170                   1175                  1180
Glu Lys Gln Asp Glu Gln Val Gly Leu Pro Gly Lys Gly Gly Lys Ala
1185                   1190                  1195                  1200
Lys Gly Lys Lys Thr Gln Met Ala Glu Val Leu Pro Ser Pro Arg Gly
                 1205                  1210                  1215
Gln Arg Val Ile Pro Arg Ile Thr Ile Glu Met Lys Ala Glu Ala Glu
             1220                  1225                  1230
Lys Lys Asn Lys Lys Lys Ile Lys Asn Glu Asn Thr Glu Gly Ser Pro
             1235                  1240                  1245
Gln Glu Asp Gly Val Glu Leu Glu Gly Leu Lys Gln Arg Leu Glu Lys
1250                   1255                  1260
Lys Gln Lys Arg Glu Pro Gly Thr Lys Thr Lys Lys Gln Thr Thr Leu
1265                   1270                  1275                  1280
Ala Phe Lys Pro Ile Lys Lys Gly Lys Lys Arg Asn Pro Trp Pro Asp
                 1285                  1290                  1295
Ser Glu Ser Asp Arg Ser Ser Asp Glu Ser Asn Phe Asp Val Pro Pro
             1300                  1305                  1310
Arg Glu Thr Glu Pro Arg Arg Ala Ala Thr Lys Thr Lys Phe Thr Met
             1315                  1320                  1325
Asp Leu Asp Ser Asp Glu Asp Phe Ser Asp Phe Asp Glu Lys Thr Asp
             1330                  1335                  1340
Asp Glu Asp Phe Val Pro Ser Asp Ala Ser Pro Pro Lys Thr Lys Thr
1345                   1350                  1355                  1360
Ser Pro Lys Leu Ser Asn Lys Glu Leu Lys Pro Gln Lys Ser Val Val
                 1365                  1370                  1375
Ser Asp Leu Glu Ala Asp Asp Val Lys Gly Ser Val Pro Leu Ser Ser
             1380                  1385                  1390
Ser Pro Pro Ala Thr His Phe Pro Asp Glu Thr Glu Ile Thr Asn Pro
             1395                  1400                  1405
Val Pro Lys Lys Asn Val Thr Val Lys Lys Thr Ala Ala Lys Ser Gln
             1410                  1415                  1420
Ser Ser Thr Ser Thr Thr Gly Ala Lys Lys Arg Ala Ala Pro Lys Gly
1425                   1430                  1435                  1440
```

Thr Lys Arg Asp Pro Ala Leu Asn Ser Gly Val Ser Gln Lys Pro Asp
              1445   1450   1455
Pro Ala Lys Thr Lys Asn Arg Arg Lys Arg Lys Pro Ser Thr Ser Asp
              1460   1465   1470
Asp Ser Asp Ser Asn Phe Glu Lys Ile Val Ser Lys Ala Val Thr Ser
         1475   1480   1485
Lys Lys Ser Lys Gly Glu Ser Asp Asp Phe His Met Asp Phe Asp Ser
      1490   1495   1500
Ala Val Ala Pro Arg Ala Lys Ser Val Arg Ala Lys Lys Pro Ile Lys
1505   1510   1515   1520
Tyr Leu Glu Glu Ser Asp Glu Asp Asp Leu Phe
              1525   1530

<210> 223
<211> 1111
<212> DNA
<213> Homo sapiens

<400> 223
ccgcgcgctc gccccgccgc tcctgctgca gccccaggcc cctcgccgcc gccaccatgg 60
acgccatcaa gaagaagatg cagatgctga agctcgacaa ggagaacgcc ttggatcgag 120
ctgagcaggc ggaggccgac aagaaggcgg cggaagacag gagcaagcag ctggaagatg 180
agctggtgtc actgcaaaag aaactcaagg gcaccgaaga tgaactggac aaatactctg 240
aggctctcaa agatgcccag gagaagctgg agctggcaga gaaaaaggcc accgatgctg 300
aagccgacgt agcttctctg aacagacgca tccagctggt tgaggaagag ttggatcgtg 360
cccaggagcg tctggcaaca gctttgcaga agctggagga agctgagaag gcagcagatg 420
agagtgagag aggcatgaaa gtcattgaga gtcgagccca aaaagatgaa gaaaaaatgg 480
aaattcagga gatccaactg aaagaggcca agcacattgc tgaagatgcc gaccgcaaat 540
acgaagaggt ggcccgtaag ctggtcatca ttgagagcga cctggaacgt gcagaggagc 600
gggctgagct ctcagaaggc aaatgtgccg agcttgaaga agaattgaaa actgtgacga 660
acaacttgaa gtcactggag gctcaggctg agaagtactc gcagaaggaa gacagatatg 720
aggaagagat caaggtcctt tccgacaagc tgaaggaggc tgagactcgg gctgagtttg 780
cggagaggtc agtaactaaa ttggagaaaa gcattgatga cttagaagac gagctgtacg 840
ctcagaaact gaagtacaaa gccatcagcg aggagctgga ccacgctctc aacgatatga 900
cttccatata gtttctttg cttcacttct cccaagactc cctcgtcgag ctggatgtcc 960
cacctctctg agctctgcat ttgtctattc tccagctgac cctggttctc tctcttagca 1020
tcctgcctta gagccaggca cacactgtgc tttctattgt acagaagctc ttcgtttcag 1080
tgtcaaataa acactgtgta agctaaaaaa a 1111

<210> 224
<211> 284
<212> PRT
<213> Homo sapiens

<400> 224
Met Asp Ala Ile Lys Lys Lys Met Gln Met Leu Lys Leu Asp Lys Glu
1   5   10   15
Asn Ala Leu Asp Arg Ala Glu Gln Ala Glu Ala Asp Lys Lys Ala Ala
              20   25   30
Glu Asp Arg Ser Lys Gln Leu Glu Asp Glu Leu Val Ser Leu Gln Lys
         35   40   45
Lys Leu Lys Gly Thr Glu Asp Glu Leu Asp Lys Tyr Ser Glu Ala Leu
      50   55   60
Lys Asp Ala Gln Glu Lys Leu Glu Leu Ala Glu Lys Lys Ala Thr Asp
65   70   75   80
Ala Glu Ala Asp Val Ala Ser Leu Asn Arg Arg Ile Gln Leu Val Glu
              85   90   95
Glu Glu Leu Asp Arg Ala Gln Glu Arg Leu Ala Thr Ala Leu Gln Lys

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                    100                     105                     110
     Leu Glu Glu Ala Glu Lys Ala Ala Asp Glu Ser Glu Arg Gly Met Lys
             115                     120                     125
     Val Ile Glu Ser Arg Ala Gln Lys Asp Glu Glu Lys Met Glu Ile Gln
         130                     135                     140
     Glu Ile Gln Leu Lys Glu Ala Lys His Ile Ala Glu Asp Ala Asp Arg
     145                     150                     155                     160
     Lys Tyr Glu Glu Val Ala Arg Lys Leu Val Ile Ile Glu Ser Asp Leu
                 165                     170                     175
     Glu Arg Ala Glu Glu Arg Ala Glu Leu Ser Glu Gly Lys Cys Ala Glu
                 180                     185                     190
     Leu Glu Glu Glu Leu Lys Thr Val Thr Asn Asn Leu Lys Ser Leu Glu
             195                     200                     205
     Ala Gln Ala Glu Lys Tyr Ser Gln Lys Glu Asp Arg Tyr Glu Glu Glu
         210                     215                     220
     Ile Lys Val Leu Ser Asp Lys Leu Lys Glu Ala Glu Thr Arg Ala Glu
     225                     230                     235                     240
     Phe Ala Glu Arg Ser Val Thr Lys Leu Glu Lys Ser Ile Asp Asp Leu
                 245                     250                     255
     Glu Asp Glu Leu Tyr Ala Gln Lys Leu Lys Tyr Lys Ala Ile Ser Glu
             260                     265                     270
     Glu Leu Asp His Ala Leu Asn Asp Met Thr Ser Ile
             275                     280
```

<210> 225
<211> 501
<212> DNA
<213> Homo sapiens

<400> 225
```
gaattcgctt tggatccatt tccatcggtc cttacagccg ctcgtcagac tccagcagcc 60
aagatggtga agcagatcga gagcaagact gcttttcagg aagccttgga cgctgcaggt 120
gataaacttg tagtagttga cttctcagcc acgtggtgtg ggccttgcaa aatgatcaac 180
cctttctttc attccctctc tgaaaagtat tccaacgtga tattccttga agtagatgtg 240
gatgactgtc aggatgttgc ttcagagtgt gaagtcaaat gcacgccaac attccagttt 300
tttaagaagg gacaaaaggt gggtgaattt tctggagcca ataaggaaaa gcttgaagcc 360
accattaatg aattagtcta atcatgtttt ctgaaaacat aaccagccat tggctattta 420
aacttgtatt tttttattta caaatataaa atatgaagac ataaccagtt gccatctgcg 480
tgacaataaa cattatgcta a                                         501
```

<210> 226
<211> 105
<212> PRT
<213> Homo sapiens

<400> 226
```
     Met Val Lys Gln Ile Glu Ser Lys Thr Ala Phe Gln Glu Ala Leu Asp
     1                   5                   10                      15
     Ala Ala Gly Asp Lys Leu Val Val Val Asp Phe Ser Ala Thr Trp Cys
                 20                      25                      30
     Gly Pro Cys Lys Met Ile Asn Pro Phe Phe His Ser Leu Ser Glu Lys
             35                      40                      45
     Tyr Ser Asn Val Ile Phe Leu Glu Val Asp Val Asp Asp Cys Gln Asp
         50                      55                      60
     Val Ala Ser Glu Cys Glu Val Lys Cys Thr Pro Thr Phe Gln Phe Phe
     65                      70                      75                      80
     Lys Lys Gly Gln Lys Val Gly Glu Phe Ser Gly Ala Asn Lys Glu Lys
                     85                      90                      95
```

```
Leu Glu Ala Thr Ile Asn Glu Leu Val
            100                 105
```

<210> 227
<211> 783
<212> DNA
<213> Homo sapiens

<400> 227
```
ggcacgagcg agttcctgtc tctctgccaa cgccgcccgg atggcttccc aaaaccgcga 60
cccagccgcc actagcgtcg ccgccgcccg taaaggagct gagccgagcg ggggcgccgc 120
ccggggtccg gtgggcaaaa ggctacagca ggagctgatg accctcatga tgtctggcga 180
taaagggatt tctgccttcc ctgaatcaga caaccttttc aaatgggtag ggaccatcca 240
tggagcagct ggaacagtat atgaagacct gaggtataag ctctcgctag agttccccag 300
tggctaccct tacaatgcgc ccacagtgaa gttcctcacg ccctgctatc accccaacgt 360
ggacacccag ggtaacatat gcctggacat cctgaaggaa aagtggtctg ccctgtatga 420
tgtcaggacc attctgctct ccatccagag ccttctagga aacccaaca ttgatagtcc 480
cttgaacaca catgctgccg agctctggaa aaaccccaca gcttttaaga agtacctgca 540
agaaacctac tcaaagcagg tcaccagcca ggagccctga cccaggctgc ccagcctgtc 600
cttgtgtcgt ctttttaatt tttccttaga tggtctgtcc tttttgtgat ttctgtatag 660
gactctttat cttgagctgt ggtatttttg ttttgttttt gtctttaaa ttaagcctcg 720
gttgagccct tgtatattaa ataaatgcat ttttgtcctt ttttaaaaaa aaaaaaaaaa 780
aaa                                                              783
```

<210> 228
<211> 179
<212> PRT
<213> Homo sapiens

<400> 228
```
Met Ala Ser Gln Asn Arg Asp Pro Ala Ala Thr Ser Val Ala Ala Ala
1               5                  10                  15
Arg Lys Gly Ala Glu Pro Ser Gly Gly Ala Ala Arg Gly Pro Val Gly
            20                  25                  30
Lys Arg Leu Gln Gln Glu Leu Met Thr Leu Met Met Ser Gly Asp Lys
        35                  40                  45
Gly Ile Ser Ala Phe Pro Glu Ser Asp Asn Leu Phe Lys Trp Val Gly
    50                  55                  60
Thr Ile His Gly Ala Ala Gly Thr Val Tyr Glu Asp Leu Arg Tyr Lys
65                  70                  75                  80
Leu Ser Leu Glu Phe Pro Ser Gly Tyr Pro Tyr Asn Ala Pro Thr Val
            85                  90                  95
Lys Phe Leu Thr Pro Cys Tyr His Pro Asn Val Asp Thr Gln Gly Asn
            100                 105                 110
Ile Cys Leu Asp Ile Leu Lys Glu Lys Trp Ser Ala Leu Tyr Asp Val
        115                 120                 125
Arg Thr Ile Leu Leu Ser Ile Gln Ser Leu Leu Gly Glu Pro Asn Ile
    130                 135                 140
Asp Ser Pro Leu Asn Thr His Ala Ala Glu Leu Trp Lys Asn Pro Thr
145                 150                 155                 160
Ala Phe Lys Lys Tyr Leu Gln Glu Thr Tyr Ser Lys Gln Val Thr Ser
            165                 170                 175
Gln Glu Pro
```

<210> 229
<211> 777

```
<212> DNA
<213> Homo sapiens

<400> 229
ggcccttgt  ctgcagagat  ggctcccaat  gcttcctgcc  tctgtgtgca  tgtccgttcc  60
gaggaatggg  atttaatgac  ctttgatgcc  aacccatatg  acagcgtgaa  aaaaatcaaa  120
gaacatgtcc  ggtctaagac  caaggttcct  gtgcaggacc  aggttctttt  gctgggctcc  180
aagatcttaa  agccacggag  aagcctctca  tcttatggca  ttgacaaaga  gaagaccatc  240
caccttaccc  tgaaagtggt  gaagcccagt  gatgaggagc  tgcccttgtt  tcttgtggag  300
tcaggtgatg  aggcaaagag  gcacctcctc  caggtgcgaa  ggtccagctc  agtggcacaa  360
gtgaaagcaa  tgatcgagac  taagacgggt  ataatccctg  agacccagat  tgtgacttgc  420
aatggaaaga  gactggaaga  tgggaagatg  atggcagatt  acggcatcag  aaagggcaac  480
ttactcttcc  tggcatctta  ttgtattgga  gggtgaccac  cctgggggatg  gggtgttggc  540
aggggtcaaa  aagcttattt  cttttaatct  cttactcaac  gaacacatct  tctgatgatt  600
tcccaaaatt  aatgagaatg  agatgagtag  agtaagattt  gggtgggatg  ggtaggatga  660
agtatattgc  ccaactctat  gtttctttga  ttctaacaca  attaattaag  tgacatgatt  720
tttactaatg  tattactgag  actagtaaat  aaatttttaa  ggcaaaatag  agcattc  777
```

```
<210> 230
<211> 165
<212> PRT
<213> Homo sapiens

<400> 230
Met Ala Pro Asn Ala Ser Cys Leu Cys Val His Val Arg Ser Glu Glu
1               5                   10                  15
Trp Asp Leu Met Thr Phe Asp Ala Asn Pro Tyr Asp Ser Val Lys Lys
            20                  25                  30
Ile Lys Glu His Val Arg Ser Lys Thr Lys Val Pro Val Gln Asp Gln
        35                  40                  45
Val Leu Leu Leu Gly Ser Lys Ile Leu Lys Pro Arg Arg Ser Leu Ser
    50                  55                  60
Ser Tyr Gly Ile Asp Lys Glu Lys Thr Ile His Leu Thr Leu Lys Val
65                  70                  75                  80
Val Lys Pro Ser Asp Glu Glu Leu Pro Leu Phe Leu Val Glu Ser Gly
                85                  90                  95
Asp Glu Ala Lys Arg His Leu Leu Gln Val Arg Arg Ser Ser Ser Val
            100                 105                 110
Ala Gln Val Lys Ala Met Ile Glu Thr Lys Thr Gly Ile Ile Pro Glu
        115                 120                 125
Thr Gln Ile Val Thr Cys Asn Gly Lys Arg Leu Glu Asp Gly Lys Met
    130                 135                 140
Met Ala Asp Tyr Gly Ile Arg Lys Gly Asn Leu Leu Phe Leu Ala Ser
145                 150                 155                 160
Tyr Cys Ile Gly Gly
                165
```

```
<210> 231
<211> 4797·
<212> DNA
<213> Homo sapiens

<400> 231
gcagtgaaca  caacctttcc  cctgagccac  tggaattgga  cagaatgccc  cattctcctc  60
tgatctccat  tcctcatgtg  tggtgtcacc  cagaagagga  ggaaagaatg  catgatgaac  120
ttctacaagc  agtatccaag  gggccggtga  tgttcaggga  tgtttccata  gacttctctc  180
aagaggaatg  ggaatgcctg  gacgctgatc  agatgaattt  atacaaagaa  gtgatgttgg  240
agaatttcag  caacctggtt  tcagtgggac  tttccaattc  taagccagct  gtgatctcct  300
```

```
tattggaaca aggaaaagag ccctggatgg ttgatagaga gctgactaga ggcctgtgtt 360
cagatctgga atcaatgtgt gagaccaaaa tattatctct aaagaagaga catttcagtc 420
aagtaataat tacccgtgaa gacatgtcta ctttttattca gcccacattt cttattccac 480
ctcaaaaaac tatgagtgaa gagaaaccat gggaatgtaa gatatgtgga aagacctta 540
atcaaaactc acaatttatc caacatcaga gaattcattt tggtgaaaaa cactatgaat 600
ctaaggagta tgggaagtcc tttagtcgtg gctcactcgt tactcgacat cagaggattc 660
acactggtaa aaaaccctat gaatgtaagg aatgtggcaa ggctttagt tgtagttcat 720
attttttctca acatcagagg attcacactg gtgagaaacc ctatgaatgt aaggaatgtg 780
gaaaagcctt taagtattgc tcaaacctta atgatcatca gagaattcac actggtgaga 840
aaccctatga atgtaaagta tgtggaaaag cctttactaa aagttcacaa cttttttctac 900
atctgagaat tcatactggt gagaaacctt atgaatgtaa agaatgtggg aaagccttta 960
ctcaacactc aaggcttatt cagcatcaga gaatgcatac tggtgagaaa ccttatgaat 1020
gtaagcagtg tgggaaggcc tttaatagtg cctcaacact tactaaccat cacagaattc 1080
atgctggtga gaagctctat gaatgtgaag aatgtagaaa ggcctttatt cagagctcag 1140
aacttattca acatcagaga atccatacag atgaaaaacc atatgaatgt aatgaatgtg 1200
ggaaggcctt taataaaggc tcaaatctta ctcgacatca gagaattcac actggtgaga 1260
aaccctatga ctgtaaggaa tgtggaaagg cttttggtag tcgctctgac ctcattcgcc 1320
atgagggaat tcatactggt tgaatgacag taaagtaaga ccattttgtt aacctttata 1380
ataattttt taaaacaggt aaggagaaca aattaggata catattatca aaggttctcc 1440
tatgtattcg tttttaaacg atacgataac aaagtaccaa gtaccaaaac cttggtggct 1500
taaaacaaga gaaatttatt ctctcatagt ttagagcctg gaaatctaaa ctcaaggtg 1560
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tgcctccatc ttcacattgc attctcgctg tgtatctctg tgtatgtctt ttatttggac 1740
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cttttaggg gatacagttc aacccataat accctgttaa aatgattttg tctaatatat 1920
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agacagagtc tcgctctgtt gcccaggctg gagtgcagtg gtgtgatctc agctcactgc 2040 ·
aacctccagc tcctgagttc aagcgattct tgtgcctcag cctctcaagt agttgggatt 2100
acaggcatgc gccaccatgc ccggctaatt ttttttttt tttttttgta tttttagtag 2160
cgacggggtt tcaccatgtt ggccaggctg gtcttgaact cctgacttca agtgatctgc 2220
ccgcctcagc ctcccaaagt gctgggatta cagacgtgag ccaccgtgat ggccaaaaca 2280 .
gactttatac caacaaaaat taaaaaggac aaagaaggtc atttataatg ataaaggata 2340
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atccataaca cagatactac tagacctaag aaaagagata gacagcaata caacaatagc 2460
aggggacttc accactccat tgacagcact agacagatca ctgggacaga aatcaacaaa 2520
gaaactctgg acttaaattg gactctacac caaatggacc caacagacat ctgaagaaca 2580
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caaaacagat acatggaatt taaacagctt gctcctgaat gatttttgga tcaatgatga 2820
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accaaaaaga tagaaaaatc tcaaatgaat agcctaacgt cacatctcaa ggaactagga 3000
aaaacaaaa caaactcaac ccaaagctgg cagaagaaaa gcaataacaa atatcagagc 3060
aggcaaaaat gagactgaga acaaaggaat gcaaaagatc aataaaagaa aaagttggtt 3120
ctttgtaaag ataaaactga cagaccacta gctagattaa ccaagaaaaa aagaagattc 3180
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tatcagcaga aactatatgc acatattaga aaacctagag gaagtggata aattcctaga 3300
aacacataac cttccaagat tgaaccaggg agaaatagga atcctcaaca gactactgag 3360
tattgaaatt gaatcagtaa tagaaaaaaa tcttgcaaaa acaaaaagcc caggaccaga 3420
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cctgatactg aagccaggca aggataaaac acacaaaaaa actacaagcc aatatccctg 3600
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aaaaagatag taacagcaca gtcaagtgga ttttattcct ggggtgtaag gatggctcaa 3720
catatgcaac tcaatacatg attcatcaca tacacagaat taaaaataag ccaggcactc 3780
acacctgtaa tcccagcact ttgcaaggcc aaggcgggca gatcacatga tgtcaagagt 3840
```

```
ttgagaccag tctggctgac atggcgaaac cctgtctcta ctaaaaatag aaaaattggc 3900
tgggcatggt ggcaggcact gtagtcccag ctacttggga ggctgaggca ggagaattac 3960
ttgaacctga gaagcggagg ttgcagtgag ctgagatagt gccattgcac tccagcctgg 4020
gcaacagagc aaattgcttg aatgtgggag gtggaggttg cagtgagccg agattatgcc 4080
attgcactcc agccggggga gcaacaaagc cagactccat ctcaaaaaaa aaccaaaaaa 4140
aatcctattt agtacaaggt acattattta ggtaatgagt ccattaaaag ccaacacttt 4200
ccccactaca ctatatgtgt atgtaacaca actgcccttg taacttccta aacctataat 4260
taagaaacaa taaaaggcaa attaagaatg ctttttttaaa aggtgggggc attatgctaa 4320
taagttactg tggatttcag agtgcagagt agaaagatca caagaattta gtgtggtagg 4380
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tactattaaa tgttttatca aactttttgat caaacaagac agcacaattt ataatttcat 4620
ttctatttct atgttatgag aaactgatca tttattcaaa tgtttaacag gcatgttcat 4680
gttactataa actcttctgt ttctccatca cgttgttggt catctttact gattacaaat 4740
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```

<210> 232
<211> 433
<212> PRT
<213> Homo sapiens

<220>
<221> VARIANT
<222> 433
<223> Xaa = Any Amino Acid

<400> 232

```
Met Pro His Ser Pro Leu Ile Ser Ile Pro His Val Trp Cys His Pro
1               5                   10                  15
Glu Glu Glu Glu Arg Met His Asp Glu Leu Leu Gln Ala Val Ser Lys
                20                  25                  30
Gly Pro Val Met Phe Arg Asp Val Ser Ile Asp Phe Ser Gln Glu Glu
            35                  40                  45
Trp Glu Cys Leu Asp Ala Asp Gln Met Asn Leu Tyr Lys Glu Val Met
        50                  55                  60
Leu Glu Asn Phe Ser Asn Leu Val Ser Val Gly Leu Ser Asn Ser Lys
65                  70                  75                  80
Pro Ala Val Ile Ser Leu Leu Glu Gln Gly Lys Glu Pro Trp Met Val
                85                  90                  95
Asp Arg Glu Leu Thr Arg Gly Leu Cys Ser Asp Leu Glu Ser Met Cys
            100                 105                 110
Glu Thr Lys Ile Leu Ser Leu Lys Lys Arg His Phe Ser Gln Val Ile
        115                 120                 125
Ile Thr Arg Glu Asp Met Ser Thr Phe Ile Gln Pro Thr Phe Leu Ile
    130                 135                 140
Pro Pro Gln Lys Thr Met Ser Glu Glu Lys Pro Trp Glu Cys Lys Ile
145                 150                 155                 160
Cys Gly Lys Thr Phe Asn Gln Asn Ser Gln Phe Ile Gln His Gln Arg
                165                 170                 175
Ile His Phe Gly Glu Lys His Tyr Glu Ser Lys Glu Tyr Gly Lys Ser
            180                 185                 190
Phe Ser Arg Gly Ser Leu Val Thr Arg His Gln Arg Ile His Thr Gly
        195                 200                 205
Lys Lys Pro Tyr Glu Cys Lys Glu Cys Gly Lys Ala Phe Ser Cys Ser
    210                 215                 220
Ser Tyr Phe Ser Gln His Gln Arg Ile His Thr Gly Glu Lys Pro Tyr
225                 230                 235                 240
Glu Cys Lys Glu Cys Gly Lys Ala Phe Lys Tyr Cys Ser Asn Leu Asn
```

```
                     245                 250                 255
Asp His Gln Arg Ile His Thr Gly Glu Lys Pro Tyr Glu Cys Lys Val
                     260                 265                 270
Cys Gly Lys Ala Phe Thr Lys Ser Ser Gln Leu Phe Leu His Leu Arg
                 275                 280                 285
Ile His Thr Gly Glu Lys Pro Tyr Glu Cys Lys Glu Cys Gly Lys Ala
             290                 295                 300
Phe Thr Gln His Ser Arg Leu Ile Gln His Gln Arg Met His Thr Gly
305                 310                 315                 320
Glu Lys Pro Tyr Glu Cys Lys Gln Cys Gly Lys Ala Phe Asn Ser Ala
                     325                 330                 335
Ser Thr Leu Thr Asn His His Arg Ile His Ala Gly Glu Lys Leu Tyr
             340                 345                 350
Glu Cys Glu Glu Cys Arg Lys Ala Phe Ile Gln Ser Ser Glu Leu Ile
         355                 360                 365
Gln His Gln Arg Ile His Thr Asp Glu Lys Pro Tyr Glu Cys Asn Glu
     370                 375                 380
Cys Gly Lys Ala Phe Asn Lys Gly Ser Asn Leu Thr Arg His Gln Arg
385                 390                 395                 400
Ile His Thr Gly Glu Lys Pro Tyr Asp Cys Lys Glu Cys Gly Lys Ala
                 405                 410                 415
Phe Gly Ser Arg Ser Asp Leu Ile Arg His Glu Gly Ile His Thr Gly
             420                 425                 430
Xaa
```

<210> 233
<211> 1860
<212> DNA
<213> Homo sapiens

<400> 233

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ggccccggcc tcgccgcccc gcgcagtacc cagcccggcc ccgccgaccc gcctctactg 120
ccggctccgc gcccttcccc gagggctgga tgatgggctg tttcgccctg caaacggtgg 180
acaccgagct gaccgcggac tcggtggagt ggtgcccgct gcaaggctgc aggcacctgc 240
tggcgtgcgg gacctaccag ctgcggcggc cggaggaccg gcctgccggc ccccagaaca 300
agggtggaat ggaagttaag gagcctcagg tccgtttagg ccgtctcttc ctgtacagtt 360
tcaatgacaa caactctatt caccctctgg tcgaggtcca agaaaagat acttctgcaa 420
tcctggacat gaaatggtgt cacatcccgg tggctggaca tgccctcttg ggcttggcag 480
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tgtattcagg gggcgacgat ggccttctga ggggctggga caccagggta cccggcaaat 840
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cccctcgtg gtcctttcct agcaacctag gaaccaagac ggcagacctg aagggtgcaa 1260
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gaccccagag tggaatgaag ccactcacag agggcatgag gaagaatggc acctggctgc 1380
aggtacagc agccaccaca cgtgactgtg gcgtgaaccc agaagaagca gactcagcct 1440
tcagcctcct ggccacctgc tccttctatg accatgcgct ccacctctgg gagtgggagg 1500
ggaactgagc ttgaaatcat gaagccccctt cccacaagga aaccaggagg gagactgcga 1560
```

```
gtgagtgccc gggaccacct catcagagat gcttactgca gccctgcagg tgcctgggca 1620
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ctgggcccct gaaagtggac tgggtgattc tgtctggcag agagtgggga aaagacgcgg 1740
tttccagctt gcagatttgt taagtttctc aggcagattt tgactttcag cctttcatac 1800
ttgtttaagc aactatttgt attaaatgaa gttttttgaa aaaaaaaaaa aaaaaaaaaa 1860
```

<210> 234
<211> 501
<212> PRT
<213> Homo sapiens

<400> 234

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Asp Pro Arg Val Arg Ala Arg Ala Asp Gly Val Pro Gly Ala Leu Arg
1               5                   10                  15
Ser Ser Gly Arg Pro Arg Pro Arg Arg Pro Ala Gln Tyr Pro Ala Arg
            20                  25                  30
Pro Arg Arg Pro Ala Ser Thr Ala Gly Ser Ala Pro Phe Pro Glu Gly
        35                  40                  45
Trp Met Met Gly Cys Phe Ala Leu Gln Thr Val Asp Thr Glu Leu Thr
    50                  55                  60
Ala Asp Ser Val Glu Trp Cys Pro Leu Gln Gly Cys Arg His Leu Leu
65                  70                  75                  80
Ala Cys Gly Thr Tyr Gln Leu Arg Arg Pro Glu Asp Arg Pro Ala Gly
                85                  90                  95
Pro Gln Asn Lys Gly Gly Met Glu Val Lys Glu Pro Gln Val Arg Leu
            100                 105                 110
Gly Arg Leu Phe Leu Tyr Ser Phe Asn Asp Asn Asn Ser Ile His Pro
        115                 120                 125
Leu Val Glu Val Gln Arg Lys Asp Thr Ser Ala Ile Leu Asp Met Lys
    130                 135                 140
Trp Cys His Ile Pro Val Ala Gly His Ala Leu Leu Gly Leu Ala Asp
145                 150                 155                 160
Ala Ser Gly Ser Ile Gln Leu Leu Arg Leu Val Glu Ser Glu Lys Ser
                165                 170                 175
His Val Leu Glu Pro Leu Ser Ser Leu Ala Leu Glu Glu Gln Cys Leu
            180                 185                 190
Ala Leu Ser Leu Asp Trp Ser Thr Gly Lys Thr Gly Arg Ala Gly Asp
        195                 200                 205
Gln Pro Leu Lys Ile Ile Ser Ser Asp Ser Thr Gly Gln Leu His Leu
    210                 215                 220
Leu Met Val Asn Glu Thr Arg Pro Arg Leu Gln Lys Val Ala Ser Trp
225                 230                 235                 240
Gln Ala His Gln Phe Glu Ala Trp Ile Ala Ala Phe Asn Tyr Trp His
                245                 250                 255
Pro Glu Ile Val Tyr Ser Gly Gly Asp Asp Gly Leu Leu Arg Gly Trp
            260                 265                 270
Asp Thr Arg Val Pro Gly Lys Phe Leu Phe Thr Ser Lys Arg His Thr
        275                 280                 285
Met Gly Val Cys Ser Ile Gln Ser Ser Pro His Arg Glu His Ile Leu
    290                 295                 300
Ala Thr Gly Ser Tyr Asp Glu His Ile Leu Leu Trp Asp Thr Arg Asn
305                 310                 315                 320
Met Lys Gln Pro Leu Ala Asp Thr Pro Val Gln Gly Gly Val Trp Arg
            325                 330                 335
Ile Lys Trp His Pro Phe His His His Leu Leu Leu Ala Ala Cys Met
        340                 345                 350
His Ser Gly Phe Lys Ile Leu Asn Cys Gln Lys Ala Met Glu Glu Arg
    355                 360                 365
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Gln	Glu	Ala	Thr	Val	Leu	Thr	Ser	His	Thr	Leu	Pro	Asp	Ser	Leu	Val
	370					375				380					

Tyr	Gly	Ala	Asp	Trp	Ser	Trp	Leu	Leu	Phe	Arg	Ser	Leu	Gln	Arg	Ala
385					390				395						400

Pro	Ser	Trp	Ser	Phe	Pro	Ser	Asn	Leu	Gly	Thr	Lys	Thr	Ala	Asp	Leu
				405					410					415	

Lys	Gly	Ala	Ser	Glu	Leu	Pro	Thr	Pro	Cys	His	Glu	Cys	Arg	Glu	Asp
			420					425				430			

Asn	Asp	Gly	Glu	Gly	His	Ala	Arg	Pro	Gln	Ser	Gly	Met	Lys	Pro	Leu
		435					440					445			

Thr	Glu	Gly	Met	Arg	Lys	Asn	Gly	Thr	Trp	Leu	Gln	Ala	Thr	Ala	Ala
	450				455					460					

Thr	Thr	Arg	Asp	Cys	Gly	Val	Asn	Pro	Glu	Glu	Ala	Asp	Ser	Ala	Phe
465					470					475					480

Ser	Leu	Leu	Ala	Thr	Cys	Ser	Phe	Tyr	Asp	His	Ala	Leu	His	Leu	Trp
				485					490					495	

Glu	Trp	Glu	Gly	Asn
			500	

<210> 235
<211> 1614
<212> DNA
<213> Homo sapiens

<400> 235
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caaccagctg actcccgtag aggaagacac tgtggaggcc agttctggag ctattgcagc 120
ctcggttgcc cggccgggga cccgagccga aaagttatcg tcagaatgtc gggcaaagac 180
cgaattgaaa tctttccctc gcgaatggca cagaccatca tgaaggctcg tttaaaggga 240
gcacagacag gtcgaaacct cctgaagaaa aaatctgatg ccttaactct tcgatttcga 300
cagatcctaa agaagataat agagactaaa atgttgatgg gcgaagtgat gagagaagct 360
gccttttcac tagctgaagc caagttcaca gcaggtgact tcagcactac agttatccaa 420
aatgtcaata aagcgcaagt gaagattcga gcgaagaaag ataatgtagc aggtgttact 480
ttgccagtat ttgaacatta ccatgaagga actgacagtt atgaactgac tggtttagcc 540
agaggtgggg aacagttggc taaattaaag aggaattatg ccaaagcagt ggaactactg 600
gtggaactag cttctctgca gacttctttt gttactttgg atgaagctat taagataacc 660
aacaggcgtg taaatgccat tgaacatgtc atcattcccc ggattgaacg tactcttgct 720
tatatcatca cagagctgga tgagagagag cgagaagagt tctataggtt aaagaaaata 780
caagagaaga aaaagattct aaaggaaaaa tctgagaagg acttggagca aaggagagca 840
gctggagagg tgttggagcc tgctaatctt ctggctgaag agaaggacga ggatcttcta 900
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tcacagtgtg taggtttgat ttgtgtggct attgattttt tggcctaaga atttcactgg 1020
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ttgcagaagc gttatcatac tccatttaca tctgtgttac acgtgatctg cttaccaagc 1200
atattaggaa atacctctta ggaagcatta gcggtctcag gccaattact gtggagcagc 1260
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tgttaccatg gtacttctca aactttgtga aaacctgcac ttttccttgc atgacaggtt 1380
cctgtcttgt ctgtcatggg agccattctg ccaatttaaa tgcgactgtg gtataaacag 1440
taaaatgatt taaaagtaag tcattccgtt tttattaatt tactgttaag tcatgttctc 1500
atgctcagat cagtagtgtc agccagagct ttctctgcag acatgtagga agtgggtagc 1560
tattttccc actccatgta ttagagtttt acaaaaaggc ttacttttga gaca 1614

<210> 236
<211> 247
<212> PRT
<213> Homo sapiens

<400> 236
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Thr Ile Met Lys Ala Arg Leu Lys Gly Ala Gln Thr Gly Arg Asn Leu
            20   25   30
Leu Lys Lys Lys Ser Asp Ala Leu Thr Leu Arg Phe Arg Gln Ile Leu
        35   40   45
Lys Lys Ile Ile Glu Thr Lys Met Leu Met Gly Glu Val Met Arg Glu
    50   55   60
Ala Ala Phe Ser Leu Ala Glu Ala Lys Phe Thr Ala Gly Asp Phe Ser
65   70   75   80
Thr Thr Val Ile Gln Asn Val Asn Lys Ala Gln Val Lys Ile Arg Ala
                85   90   95
Lys Lys Asp Asn Val Ala Gly Val Thr Leu Pro Val Phe Glu His Tyr
            100   105   110
His Glu Gly Thr Asp Ser Tyr Glu Leu Thr Gly Leu Ala Arg Gly Gly
        115   120   125
Glu Gln Leu Ala Lys Leu Lys Arg Asn Tyr Ala Lys Ala Val Glu Leu
    130   135   140
Leu Val Glu Leu Ala Ser Leu Gln Thr Ser Phe Val Thr Leu Asp Glu
145   150   155   160
Ala Ile Lys Ile Thr Asn Arg Arg Val Asn Ala Ile Glu His Val Ile
                165   170   175
Ile Pro Arg Ile Glu Arg Thr Leu Ala Tyr Ile Ile Thr Glu Leu Asp
            180   185   190
Glu Arg Glu Arg Glu Glu Phe Tyr Arg Leu Lys Lys Ile Gln Glu Lys
        195   200   205
Lys Lys Ile Leu Lys Glu Lys Ser Glu Lys Asp Leu Glu Gln Arg Arg
    210   215   220
Ala Ala Gly Glu Val Leu Glu Pro Ala Asn Leu Leu Ala Glu Glu Lys
225   230   235   240
Asp Glu Asp Leu Leu Phe Glu
            245

<210> 237
<211> 1658
<212> DNA
<213> Homo sapiens

<400> 237
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gaactgccag ccaagatcct ggttgagttt gtggtggact ctcagaagaa agacaagctg 180
ctctgcagcc agcttcaggt agcggatttc ctgcagaaca tcctggctca ggaggacact 240
gctaagggtc tcgacccctt ggcttctgaa gacacgagcc gacagaaggc aattgcagct 300
aaggaacaat ggaaagagct gaaggccacc tacagggagc acgtagaggc catcaaaatt 360
ggcctcacca aggccctgac tcagatggag gaagcccaga ggaaacggac acaactccgg 420
gaagcctttg agcagctcca ggccaagaaa caaatggcca tggagaaacg cagagcagtc 480
cagaaccagt ggcagctaca acaggagaag catctgcagc atctggcgga ggtttctgca 540
gaggtgaggg agcgtaagac agggactcag caggagcttg acggggtgtt tcagaaactt 600
ggaaacctga agcagcaggc agaacaggag cgggacaagc tgcagaggta tcagaccttc 660
ctccagcttc tgtataccct gcagggtaag ctgttgttcc ctgaggctga ggctgaggca 720
gagaatcttc cagatgataa accccagcag ccgactcgac cccaggagca gagtacagga 780
gacaccatgg ggagagaccc tggtgtgtcc ttcaaggctg ttggtctaca acctgctgga 840
gatgtaaatt tgccatgact tcctggagga cagcagcatg gagaaagatc ctagaaaagg 900
cctctgactt ccctcacctc ccaaccatca ttacaggaaa gactgtgaac tcctgagttc 960
agcttgattt ctgactacat cccagcaagc tctggcatct gtggattaaa atccctggat 1020
ctctctcagt tgtgtatttg ttcatcttca tatgctggca ggaacaacta ttaatacaga 1080

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tactcagaag ccaataacat gacaggagct gggactggtt tgaacacagg gtgtgcagat 1140
ggggagggg  tactggcctt gggcctccta tgatgcagac atggtgaatt taattcaagg 1200
aggaggagaa tgttttaggc aggtggttat atgtgggaag ataattttat tcatggatcc 1260
aaatgtttgt tgagtccttt ctttgtgcta aggttcttgc ggtgaaccag aattataaca 1320
gtgagctcat ctgactgttt taggatgtac agcctagtgt taacattctt ggtatctttt 1380
tgtgccttat ctaaaacatt tctcgatcac tggtttcaga tgttcattta ttatattctt 1440
ttcaaagatt cagagattgg cttttgtcat ccactattgt atgttttgtt tcattgacct 1500
ctagtgatac cttgatcttt cccactttct gttttcggat tggagaagat gtaccttttt 1560
tgtcaactct tactttttatc agatgatcaa ctcacgtatt tggatcttta tttgttttct 1620
caaataaata tttaaggtta aaaaaaaaaa aaaaaaa                           1658
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<210> 238
<211> 277
<212> PRT
<213> Homo sapiens

<400> 238

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1               5                   10                  15
Ala Glu Val Ala Gly Ile Leu Glu Pro Val Gly Leu Gln Glu Glu Ala
            20                  25                  30
Glu Leu Pro Ala Lys Ile Leu Val Glu Phe Val Val Asp Ser Gln Lys
        35                  40                  45
Lys Asp Lys Leu Leu Cys Ser Gln Leu Gln Val Ala Asp Phe Leu Gln
    50                  55                  60
Asn Ile Leu Ala Gln Glu Asp Thr Ala Lys Gly Leu Asp Pro Leu Ala
65                  70                  75                  80
Ser Glu Asp Thr Ser Arg Gln Lys Ala Ile Ala Ala Lys Glu Gln Trp
                85                  90                  95
Lys Glu Leu Lys Ala Thr Tyr Arg Glu His Val Glu Ala Ile Lys Ile
            100                 105                 110
Gly Leu Thr Lys Ala Leu Thr Gln Met Glu Glu Ala Gln Arg Lys Arg
        115                 120                 125
Thr Gln Leu Arg Glu Ala Phe Glu Gln Leu Gln Ala Lys Lys Gln Met
        130                 135                 140
Ala Met Glu Lys Arg Arg Ala Val Gln Asn Gln Trp Gln Leu Gln Gln
145                 150                 155                 160
Glu Lys His Leu Gln His Leu Ala Glu Val Ser Ala Glu Val Arg Glu
                165                 170                 175
Arg Lys Thr Gly Thr Gln Gln Glu Leu Asp Gly Val Phe Gln Lys Leu
            180                 185                 190
Gly Asn Leu Lys Gln Gln Ala Glu Gln Glu Arg Asp Lys Leu Gln Arg
        195                 200                 205
Tyr Gln Thr Phe Leu Gln Leu Leu Tyr Thr Leu Gln Gly Lys Leu Leu
    210                 215                 220
Phe Pro Glu Ala Glu Ala Glu Ala Glu Asn Leu Pro Asp Asp Lys Pro
225                 230                 235                 240
Gln Gln Pro Thr Arg Pro Gln Glu Gln Ser Thr Gly Asp Thr Met Gly
                245                 250                 255
Arg Asp Pro Gly Val Ser Phe Lys Ala Val Gly Leu Gln Pro Ala Gly
            260                 265                 270
Asp Val Asn Leu Pro
            275
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